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spectra  
NEWS 4 MAR 31 CA/Cplus and CASREACT patent number format for U.S.  
applications updated  
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NEWS 6 MAR 31 EMBASE, EMBAL, and LEMBASE reloaded with enhancements  
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predefined hit display formats  
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NEWS 10 APR 28 IMSRESEARCH reloaded with enhancements  
NEWS 11 MAY 30 INPAFAMDB now available on STN for patent family  
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NEWS 12 MAY 30 DGENE, PCTGEN, and USGENE enhanced with new homology  
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NEWS 13 JUN 06 EPFULL enhanced with 260,000 English abstracts  
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Assistant and BLAST plug-in  
NEWS 21 JUN 30 STN AnaVist enhanced with database content from EPFULL  
NEWS 22 JUL 28 CA/Cplus patent coverage enhanced  
NEWS 23 JUL 28 EPFULL enhanced with additional legal status  
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NEWS 25 JUL 28 STN Viewer performance improved  
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10541328

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STRUCTURE FILE UPDATES: 2 AUG 2008 HIGHEST RN 1037774-47-2

conducting SmartSELECT searches.

predicted properties as well as tags indicating availability of

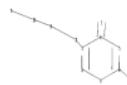
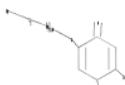
10541328

experimental property data in the original document. For information on property searching in REGISTRY, refer to:

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=>

Uploading C:\Program Files\Stnexp\Queries\10541328.str



chain nodes :

7 9 10 12 15 16

ring nodes :

1 2 3 4 5 6

chain bonds :

3-12 4-7 6-16 9-10 9-12 10-15

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

1-2 1-6 2-3 3-4 3-12 4-5 4-7 5-6 6-16 9-10 9-12 10-15

isolated ring systems :

containing 1 :

G1:O,S,N

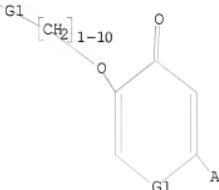
10541328

Match level :  
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 9:CLASS 10:Atom 12:CLASS  
15:Atom 16:CLASS

L1 STRUCTURE UPLOADED

=> d 11  
L1 HAS NO ANSWERS  
L1 STR

Hy



G1 O, S, N

Structure attributes must be viewed using STN Express query preparation.

=> s 11  
SAMPLE SEARCH INITIATED 10:41:50 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 27269 TO ITERATE

7.3% PROCESSED 2000 ITERATIONS  
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED) 0 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 535500 TO 555260  
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s 11 sss full  
FULL SEARCH INITIATED 10:41:57 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 553581 TO ITERATE

100.0% PROCESSED 553581 ITERATIONS 69 ANSWERS  
SEARCH TIME: 00.00.08

L3 69 SEA SSS FUL L1

=> FIL HCAPLUS			
COST IN U.S. DOLLARS		SINCE FILE	TOTAL
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		178.36	178.57

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FILE COVERS 1907 - 4 Aug 2008 VOL 149 ISS 6  
 FILE LAST UPDATED: 3 Aug 2008 (20080803/ED)

HCplus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2008.

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This file contains CAS Registry Numbers for easy and accurate substance identification.

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L4      5 L3

=> s 14 and py<=2003
      24005635 PY<=2003
L5      1 L4 AND PY<=2003

=> d 14 ibib abs hitstr tot
```

L4 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 2008:622524 HCAPLUS  
 TITLE: Characterization of EHT 1864, a novel small molecule inhibitor of Rac family small GTPases  
 AUTHOR(S): Onesto, Cercina; Shutes, Adam; Picard, Virginie; Schweighoffer, Fabien; Der, Channing J.  
 CORPORATE SOURCE: Lineberger Comprehensive Cancer Center, Department of Pharmacology, University of North Carolina at Chapel Hill, Chapel Hill, NC, USA  
 SOURCE: Methods in Enzymology (2008), 439(Small GTPases in Disease, Part B), 111-129  
 CODEN: MENZAU; ISSN: 0076-6879  
 PUBLISHER: Elsevier  
 DOCUMENT TYPE: Journal; General Review  
 LANGUAGE: English  
 AB A review. There is now considerable exptl. evidence that aberrant

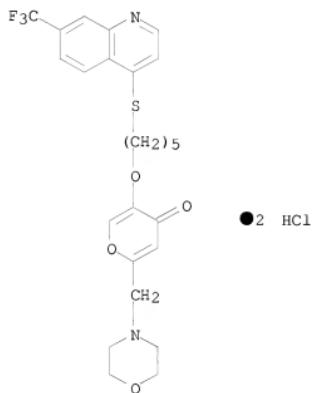
activation of Rho family small GTPases promotes uncontrolled proliferation, invasion, and metastatic properties of human cancer cells. Therefore, there is considerable interest in the development of small mol. inhibitors of Rho GTPase function. However, to date, most efforts have focused on inhibitors that block Rho GTPase function indirectly, either by targeting enzymes involved in post-translational processing or downstream protein kinase effectors. We have reported the identification and characterization of the EHT 1864 small mol. as an inhibitor of Rac family small GTPases, placing Rac1 in an inert and inactive state and then impairing Rac1-mediated functions *in vivo*. Our work suggests that EHT 1864 selectively inhibits Rac1 downstream signaling and cellular transformation by a novel mechanism involving guanine nucleotide displacement. This chapter provides the details for some of the biochem. and biol. methods used to characterize the mode of action of EHT 1864 on Rac1 and its impact on Rac1-dependent cellular functions.

IT 754240-09-0, EHT1864

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(biochem. and biol. methods may be useful to characterize Rho GTPase specificity and mechanism of action of EHT 1864 on Rac1 and its impact on Rac1-dependent cellular function in mouse)

RN 754240-09-0 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:2) (CA INDEX NAME)



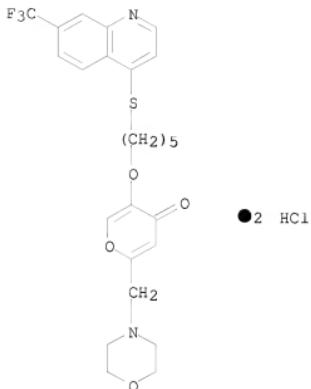
L4 ANSWER 2 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:1368320 HCPLUS

DOCUMENT NUMBER: 148:232138

TITLE: Specificity and Mechanism of Action of EHT 1864, a Novel Small Molecule Inhibitor of Rac Family Small GTPases

AUTHOR(S): Shutes, Adam; Onesto, Cercina; Picard, Virginie;  
 Leblond, Bertrand; Schweighoffer, Fabien; Der,  
 Channing J.  
 CORPORATE SOURCE: Lineberger Comprehensive Cancer Center, University of  
 North Carolina, Chapel Hill, NC, 27599, USA  
 SOURCE: Journal of Biological Chemistry (2007), 282(49),  
 35666-35678  
 CODEN: JBCHA3; ISSN: 0021-9258  
 PUBLISHER: American Society for Biochemistry and Molecular  
 Biology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB There is now considerable exptl. evidence that aberrant activation of Rho  
 family small GTPases promotes the uncontrolled proliferation, invasion,  
 and metastatic properties of human cancer cells. Therefore, there is  
 considerable interest in the development of small mol. inhibitors of Rho  
 GTPase function. However, to date, most efforts have focused on  
 inhibitors that indirectly block Rho GTPase function, by targeting either  
 enzymes involved in post-translational processing or downstream protein  
 kinase effectors. We recently determined that the EHT 1864 small mol. can  
 inhibit Rac function *in vivo*. In this study, we evaluated the biol. and  
 biochem. specificities and biochem. mechanism of action of EHT 1864. We  
 determined that EHT 1864 specifically inhibited Rac1-dependent platelet-derived  
 growth factor-induced lamellipodia formation. Furthermore, our biochem.  
 analyses with recombinant Rac proteins found that EHT 1864 possesses high  
 affinity binding to Rac1, as well as the related Rac1b, Rac2, and Rac3  
 isoforms, and this association promoted the loss of bound nucleotide,  
 inhibiting both guanine nucleotide association and Tiam1 Rac guanine  
 nucleotide exchange factor-stimulated exchange factor activity *in vitro*.  
 EHT 1864 therefore places Rac in an inert and inactive state, preventing  
 its engagement with downstream effectors. Finally, we evaluated the  
 ability of EHT 1864 to block Rac-dependent growth transformation, and we  
 determined that EHT 1864 potently blocked transformation caused by  
 constitutively activated Rac1, as well as Rac-dependent transformation  
 caused by Tiam1 or Ras. Taken together, our results suggest that EHT 1864  
 selectively inhibits Rac downstream signaling and transformation by a  
 novel mechanism involving guanine nucleotide displacement.  
 IT 754240-09-0, EHT 1864  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (EHT 1864 specifically inhibits Rac1-dependent platelet-derived growth  
 factor-induced lamellipodia formation)  
 RN 754240-09-0 HCPLUS  
 CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-  
 quinolinyl]thiopentyl)oxyl]-, hydrochloride (1:2) (CA INDEX NAME)



REFERENCE COUNT: 61 THERE ARE 61 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 2005:1191062 HCAPLUS  
 DOCUMENT NUMBER: 144:68139  
 TITLE: RAC1 Inhibition Targets Amyloid Precursor Protein Processing by  $\gamma$ -Secretase and Decreases  $\text{A}\beta$  Production in Vitro and in Vivo  
 AUTHOR(S): Desire, Laurent; Bourdin, Jerome; Loiseau, Nadia; Peillon, Helene; Picard, Virginie; De Oliveira, Catherine; Bachelot, Florence; Leblond, Bertrand; Taverne, Thierry; Beausoleil, Eric; Lacombe, Sandrine; Drouin, Dominique; Schweighoffer, Fabien  
 CORPORATE SOURCE: Exonhit Therapeutics, Paris, 75013, Fr.  
 SOURCE: Journal of Biological Chemistry (2005), 280(45), 37516-37525  
 PUBLISHER: American Society for Biochemistry and Molecular Biology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB  $\beta$ -Amyloid peptides ( $\text{A}\beta$ ) that form the senile plaques of Alzheimer disease consist mainly of 40- and 42-amino acid ( $\text{A}\beta$  40 and  $\text{A}\beta$  42) peptides generated from the cleavage of the amyloid precursor protein (APP). Generation of  $\text{A}\beta$  involves  $\beta$ -secretase and  $\gamma$ -secretase activities and is regulated by membrane trafficking of the proteins involved in  $\text{A}\beta$  production. Here we describe a new small mol., EHT 1864, which blocks the Rac1 signaling pathways. In vitro, EHT 1864 blocks  $\text{A}\beta$  40 and  $\text{A}\beta$  42 production but does not impact sAPP $\alpha$  levels and does not inhibit  $\beta$ -secretase. Rather, EHT 1864 modulates APP processing at the level of  $\gamma$ -secretase to prevent

A $\beta$  40 and A $\beta$  42 generation. This effect does not result from a direct inhibition of the  $\gamma$ -secretase activity and is specific for APP cleavage, since EHT 1864 does not affect Notch cleavage. In vivo, EHT 1864 significantly reduces A $\beta$  40 and A $\beta$  42 levels in guinea pig brains at a threshold that is compatible with delaying plaque accumulation and/or clearing the existing plaque in brain. EHT 1864 is the first derivative of a new chemical series that consists of candidates for inhibiting A $\beta$  formation in the brain of AD patients. Our findings represent the first pharmacol. validation of Rac1 signaling as a target for developing novel therapies for Alzheimer disease.

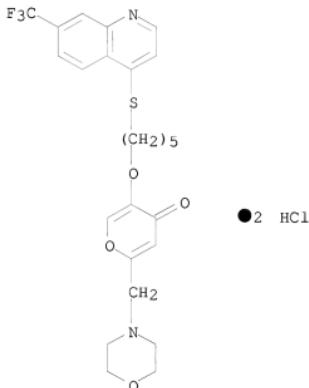
IT 754240-09-0

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(EHT 1864; EHT 1864 blocked A $\beta$ 1-40 and A $\beta$ 1-42 production)

RN 754240-09-0 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:2) (CA INDEX NAME)



REFERENCE COUNT:

55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:740320 HCPLUS

DOCUMENT NUMBER: 141:260557

TITLE: Preparation of novel antiproliferative and

antiangiogenic agents, in particular  
quinoline-derivatized pyranones, for treating cell  
proliferative diseases, retinopathies and arthritis  
Leblond, Bertrand; Petit, Silvere; Picard, Virginie;  
Taverne, Thierry; Schweighoffer, Fabien

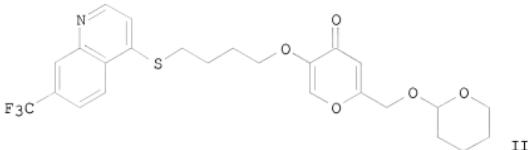
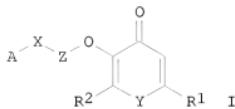
INVENTOR(S):

PATENT ASSIGNEE(S): Exonhit Therapeutics Sa, Fr.

SOURCE: PCT Int. Appl., 156 pp.

CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004076445	A2	20040910	WO 2004-IB926	20040227
WO 2004076445	A3	20050106		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
EP 1471063	A1	20041027	EP 2003-290490	20030228
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AU 2004215577	A1	20040910	AU 2004-215577	20040227
CA 2516239	A1	20040910	CA 2004-2516239	20040227
EP 1597253	A2	20051123	EP 2004-715422	20040227
EP 1597253	B1	20060809		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
CN 1747952	A	20060315	CN 2004-80003820	20040227
JP 2006519221	T	20060824	JP 2006-502497	20040227
AT 335734	T	20060915	AT 2004-715422	20040227
US 20060183749	A1	20060817	US 2005-541328	20050830
PRIORITY APPLN. INFO.:			EP 2003-290490	A 20030228
			WO 2004-IB926	W 20040227
OTHER SOURCE(S):	MARPAT 141:260557			
GI				



AB Title compds. I [wherein R1 = [(tetrahydropyran-2-yl)oxy]methyl, CH2-B, (morpholin-4-yl)methyl, pyrrolidin-1-ylmethyl, etc.; B = halo, OH, OCH2OMe, OCH2OCH2CH2OMe, OSO2-alkyl, OTBDMS; R2 = H, alk(en)yl; X, Y = independently O, S, NH and derivs.; A = quinolin-4-yl, quinolin-8-yl, benzo[b]thiophen-7-yl, quinazolin-4-yl; Z = (CH2)n, optionally interrupted by a heteroatom, C(:O) or aryldialkyl, especially xlenyl, group; n = 1-10; their tautomers, optical and geometrical isomers, racemates, salts, hydrates and mixts.] were prepared as antiproliferative agents and angiogenesis inhibitors. Nine biol. assays are given. For example, II was prepared, in 2 steps, from pyranone III, 1,4-dibromobutane, and 7-(trifluoromethyl)-4-quinolinethiol. In an *in vitro* cell viability assay, selected I showed an IC50 < 4  $\mu$ M and < 9  $\mu$ M against HCT116 and MDA-MB-231 tumoral cell lines, demonstrating their cytostatic mode of action. I are useful for treating various diseases associated with abnormal cell proliferation, including cancer, especially leukemia, or associated with unregulated angiogenesis including growth and metastasis of solid tumors, ocular diseases, especially retinopathies, or arthritis.

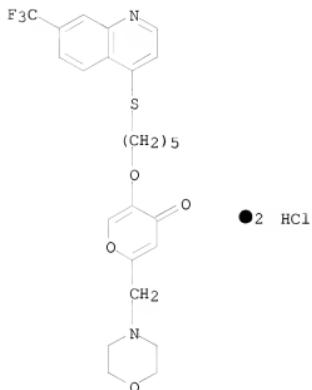
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754240-17-0P, 5-[[5-[(7-(Trifluoromethyl)quinolin-4-ylthio)pentyl]oxy]-2-[(4-methylpiperazin-1-yl)methyl]-4H-pyran-4-one trihydrochloride 754240-19-2P, 5-[[5-[(7-(Trifluoromethyl)quinolin-4-ylthio)pentyl]oxy]-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one trihydrochloride

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

RN 754240-09-0 HCAPLUS

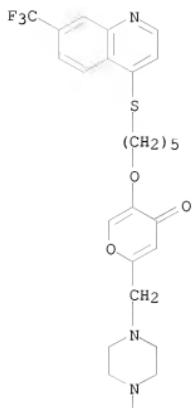
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[5-[7-(trifluoromethyl)-4-

quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:2) (CA INDEX NAME)



RN 754240-17-0 HCPLUS

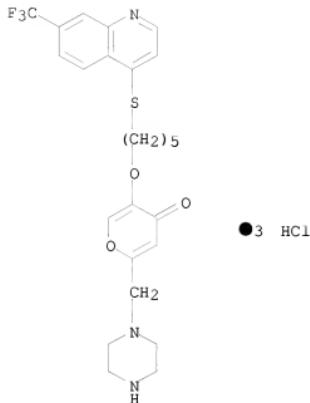
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:3) (CA INDEX NAME)



● 3 HCl

RN 754240-19-2 HCPLUS

CN 4H-Pyran-4-one, 2-(1-piperazinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]-, hydrochloride (1:3) (CA INDEX NAME)



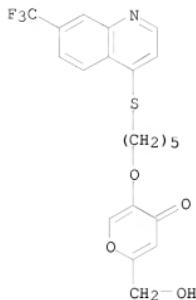
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RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

RN 754239-59-3 HCPLUS

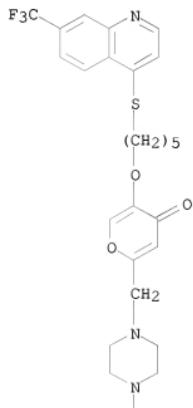
CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[7-(trifluoromethyl)-4-quinolinyl]thiopentyl)oxy]- (CA INDEX NAME)



RN 754240-15-8 HCPLUS

CN 1-Piperazinecarboxylic acid, 4-[(4-oxo-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl)thiolpentyl]oxy)-4H-pyran-2-yl]methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

PAGE 1-A



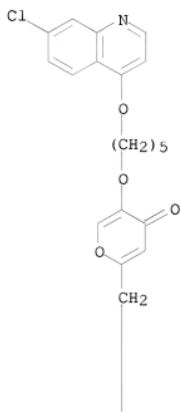
PAGE 2-A



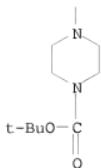
RN 754240-16-9 HCPLUS

CN 1-Piperazinecarboxylic acid, 4-[[5-[[5-[(7-chloro-4-quinolinyl)oxypentyl]oxy]-4-oxo-4H-pyran-2-yl]methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



IT 754239-47-9P, 5-[[5-((6-Fluoro-2-methylquinolin-4-yloxy)pentyl)oxy]-2-[(tetrahydropyran-2-yl)oxy]methyl]-4H-pyran-4-one 754239-48-0P

$$\begin{aligned}
 & 5-[[5-(6-Fluoro-2-trifluoromethylquinolin-4-yloxy)pentyl]oxy]-2- \\
 & [[(tetrahydropyran-2-yl)oxy]methyl]-4H-pyran-4-one 754239-49-1P, \\
 & 5-[[5-(7-Propylquinolin-8-yloxy)pentyl]oxy]-2-[[tetrahydropyran-2- \\
 & yl)oxy]methyl]-4H-pyran-4-one 754239-50-4P, 5-[[ \\
 & (Benz[b]thiophen-7-yl)oxy]pentyl]oxy]-2-[[tetrahydropyran-2- \\
 & yl)oxy]methyl]-4H-pyran-4-one 754239-51-5P, 2-[[Tetrahydropyran- \\
 & 2-yl)oxy]methyl]-5-[[5-(7-(trifluoromethyl)quinolin-4- \\
 & yl)sulfanyl]pentyl]oxy]-4H-pyran-4-one 754239-53-7P, \\
 & 2-[[Tetrahydropyran-2-yl)oxy]methyl]-5-[[4-[[7-(trifluoromethyl)quinolin-4- \\
 & yl)sulfanyl]butoxy]-4H-pyran-4-one 754239-55-9P, \\
 & 2-[[Tetrahydropyran-2-yl)oxy]methyl]-5-[[6-[[7-(trifluoromethyl)quinolin-4- \\
 & yl)sulfanyl]hexyloxy]-4H-pyran-4-one 754239-56-0P, \\
 & 2-Hydroxymethyl-5-[[5-(7-trifluoromethylquinolin-4-ylsulfanyl)pentyl]oxy]- \\
 & 4H-pyran-4-one hydrochloride 754239-60-6P, 2- \\
 & [(Methoxymethoxy)methyl]-5-[[5-(7-(trifluoromethyl)quinolin-4- \\
 & ylsulfanyl)pentyl]oxy]-4H-pyran-4-one 754239-62-8P, \\
 & 2-Chloromethyl-5-[[5-(7-trifluoromethylquinolin-4-ylsulfanyl)pentyl]oxy]- \\
 & 4H-pyran-4-one 754239-63-9P, 2-(4-Methylpiperazin-1-ylmethyl)-5- \\
 & [[5-[[7-(trifluoromethyl)quinolin-4-yl]sulfanyl]pentyl]oxy]-4H-pyran-4-one \\
 & 754239-64-0P, 2-[(Morpholin-4-yl)methyl]-5-[[5-[[7- \\
 & (trifluoromethyl)quinolin-4-yl]sulfanyl]pentyl]oxy]-4H-pyran-4-one \\
 & 754239-66-2P, 5-[[7-[[7-(Trifluoromethyl)quinolin-4- \\
 & ylthio]heptyl]oxy]-2-[[tetrahydro-2H-pyran-2-yl)oxy]methyl]-4H-pyran-4- \\
 & one 754239-68-4P, 5-[[18-[[7-(Trifluoromethyl)quinolin-4- \\
 & ylthio]octyl]oxy]-2-[[tetrahydro-2H-pyran-2-yl)oxy]methyl]-4H-pyran-4-one \\
 & 754239-80-0P, 5-[[15-[[7-(Trifluoromethyl)quinolin-4- \\
 & ylthiopentyl]oxy]-2-[(piperidin-1-yl)methyl]-4H-pyran-4-one \\
 & 754239-81-1P, 5-[[5-[[7-(Trifluoromethyl)quinolin-4- \\
 & ylthiopentyl]oxy]-2-(thiomorpholinomethyl)-4H-pyran-4-one \\
 & 754239-82-2P, 2-[(Diethylamino)methyl]-5-[[5-[[7- \\
 & (trifluoromethyl)quinolin-4-ylthio]pentyl]oxy]-4H-pyran-4-one \\
 & 754239-83-3P, 5-[[5-[(6-(Trifluoromethyl)quinolin-4- \\
 & yloxy]pentyl]oxy]-2-(morpholinomethyl)-4H-pyran-4-one 754239-84-4P \\
 & , 5-[[5-[(7-(Trifluoromethyl)quinolin-4-yloxy]pentyl]oxy]-2-[(4- \\
 & methylpiperazin-1-yl)methyl]-4H-pyran-4-one 754239-85-5P, \\
 & 5-[[5-[[7-(Trifluoromethyl)quinolin-4-yloxy]pentyl]oxy]-2- \\
 & (morpholinomethyl)-4H-pyran-4-one 754239-86-6P, \\
 & 4-[[5-[[6-(4-Methylpiperazin-1-ylmethyl)-4-oxo-4H-pyran-3- \\
 & yl]oxy]pentyl]oxy]-7-trifluoromethylquinoline-3-carboxylic acid ethyl \\
 & ester 754239-87-7P, 4-[[5-[[6-(Morpholin-4-yl)methyl]-4-oxo-4H- \\
 & pyran-3-yl]oxy]pentyl]oxy]-7-trifluoromethylquinoline-3-carboxylic acid \\
 & ethyl ester 754239-88-8P, 5-[[5-[(Trifluoromethyl)quinolin-4- \\
 & yloxy]pentyl]oxy]-2-(4-methylpiperazin-1-yl)methyl]-4H-pyran-4-one \\
 & 754239-89-9P, 5-[[5-[(Trifluoromethyl)quinolin-4- \\
 & yloxy]pentyl]oxy]-2-(morpholinomethyl)-4H-pyran-4-one 754239-90-2P \\
 & , 5-[[5-(Quinazolin-4-yloxy)pentyl]oxy]-2-[(pyrrolidin-1-yl)methyl]-4H- \\
 & pyran-4-one 754239-91-3P, 5-[[5-(Quinazolin-4-yloxy)pentyl]oxy]- \\
 & 2-(morpholinomethyl)-4H-pyran-4-one 754239-92-4P, \\
 & 5-[[5-(Quinazolin-4-yloxy)pentyl]oxy]-2-[(4-methylpiperazin-1-yl)methyl]- \\
 & 4H-pyran-4-one 754239-93-5P, 5-[[5-[(Trifluoromethyl)quinolin- \\
 & 4-yloxy]pentyl]oxy]-2-(fluoromethyl)-4H-pyran-4-one 754240-07-8P \\
 & , 5-[[2-[[7-(Trifluoromethyl)quinolin-4-ythio]ethoxy]-2-[(tetrahydro-2H- \\
 & pyran-2-yl)oxy]methyl]-4H-pyran-4-one 754240-08-9P, \\
 & 5-[[2-[[7-(Trifluoromethyl)quinolin-4-yloxy]ethoxy]-2-(morpholinomethyl)-4H- \\
 & pyran-4-one 754240-18-1P, 5-[[5-(7-Chloroquinolin-4- \\
 & yloxy)pentyl]oxy]-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one \\
 & trihydrochloride 754240-21-6P, 5-[[5-[[7- \\
 \end{aligned}$$

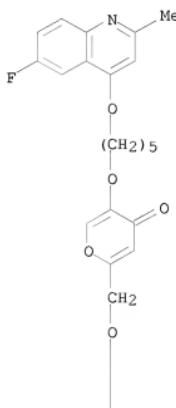
(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy]-2-[(4-acetylpirazin-1-yl)methyl]-4H-pyran-4-one 754240-22-7P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N,N-diethylpiperazine-1-carboxamide 754240-23-8P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-2-[(4-(pivaloyl)piperazin-1-yl)methyl]-4H-pyran-4-one 754240-24-9P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N,N-diisopropylpiperazine-1-carboxamide 754240-25-0P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-2-[(4-Methylsulfonylpiperazin-1-yl)methyl]-4H-pyran-4-one 754240-26-1P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N-tert-butylpiperazine-1-carboxamide 754240-27-2P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N-methylpiperazine-1-carboxamide 754240-28-3P, 5-[(5-[(7-Chloroquinolin-4-yloxy)pentyl]oxy)-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

RN 754239-47-9 HCAPLUS

CN 4H-Pyran-4-one, 5-[(5-[(6-fluoro-2-methyl-4-quinolinyl)oxy]pentyl]oxy)-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl]- (CA INDEX NAME)

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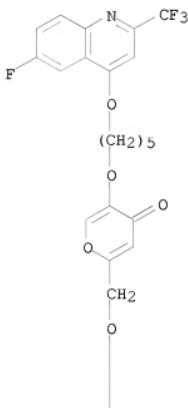


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RN 754239-48-0 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[(5-[(6-fluoro-2-(trifluoromethyl)-4-quinolinyl)oxy]pentyl)oxy]-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl- (CA INDEX NAME)

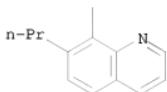
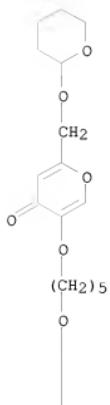
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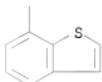
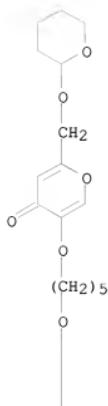
PAGE 2-A



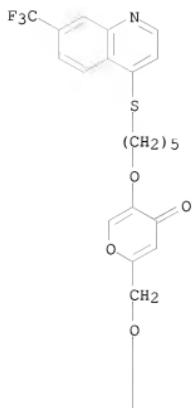
RN 754239-49-1 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[(5-[(7-propyl-8-quinolinyl)oxy]pentyl)oxy]-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl- (CA INDEX NAME)



RN 754239-50-4 HCPLUS  
CN 4H-Pyran-4-one, 5-[[5-(benzo[b]thien-7-yloxy)pentyl]oxy]-2-[[((tetrahydro-2H-pyran-2-yl)oxy)methyl]- (CA INDEX NAME)

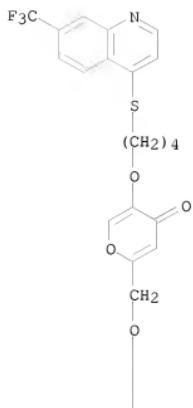


RN 754239-51-5 HCPLUS  
CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl)thio]pentyl)oxy]- (CA INDEX NAME)



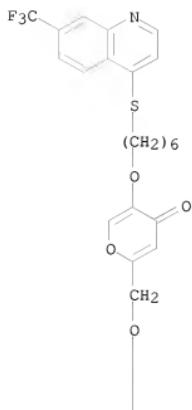
RN 754239-53-7 HCPLUS

CN 4H-Pyran-4-one, 2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[4-[(7-(trifluoromethyl)-4-quinolinyl)thio]butoxy]- (CA INDEX NAME)



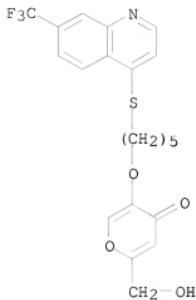
RN 754239-55-9 HCPLUS

CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(6-[(7-(trifluoromethyl)-4-quinoliny]thio]hexyl]oxy]- (CA INDEX NAME)



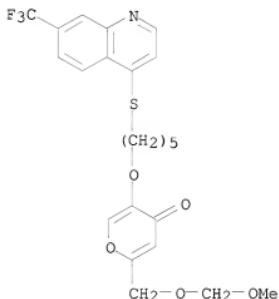
RN 754239-56-0 HCPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy]-, hydrochloride (1:1) (CA INDEX NAME)

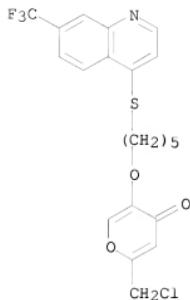


● HCl1

RN 754239-60-6 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(methoxymethoxy)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



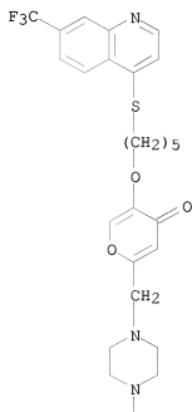
RN 754239-62-8 HCAPLUS  
 CN 4H-Pyran-4-one, 2-(chloromethyl)-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



RN 754239-63-9 HCPLUS

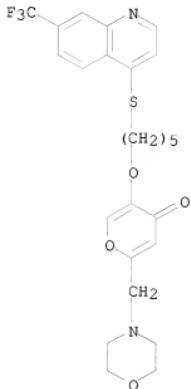
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy]- (CA INDEX NAME)

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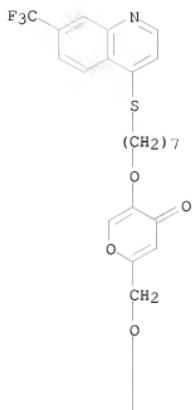




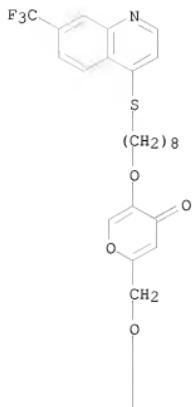
RN 754239-64-0 HCAPLUS  
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinoliny]thio)pentyl]oxy)- (CA INDEX NAME)



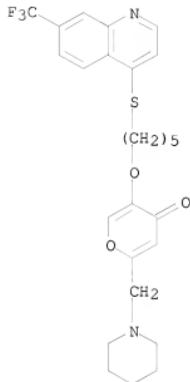
RN 754239-66-2 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(7-[(7-(trifluoromethyl)-4-quinoliny]thio)heptyl]oxy)- (CA INDEX NAME)



RN 754239-68-4 HCPLUS  
CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(8-[(7-(trifluoromethyl)-4-quinoliny]thio]octyl)oxy]- (CA INDEX NAME)

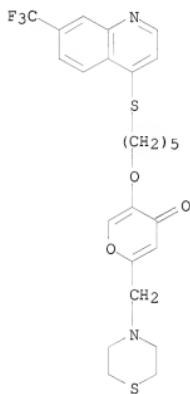


RN 754239-80-0 HCPLUS  
CN 4H-Pyran-4-one, 2-(1-piperidinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy]- (CA INDEX NAME)



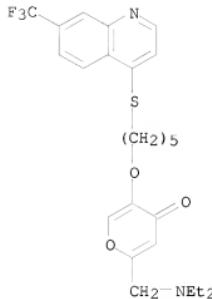
RN 754239-81-1 HCPLUS

CN 4H-Pyran-4-one, 2-(4-thiomorpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy)- (CA INDEX NAME)



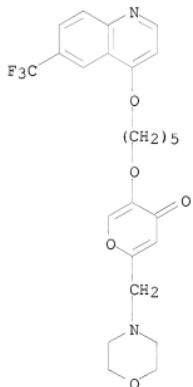
RN 754239-82-2 HCPLUS

CN 4H-Pyran-4-one, 2-[(diethylamino)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy)- (CA INDEX NAME)



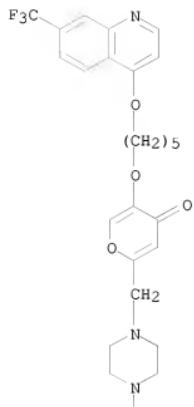
RN 754239-83-3 HCAPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[[5-[[6-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



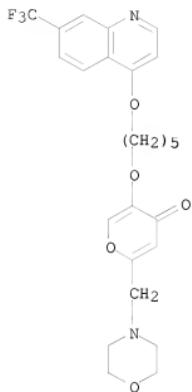
RN 754239-84-4 HCAPLUS

CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



RN 754239-85-5 HCPLUS

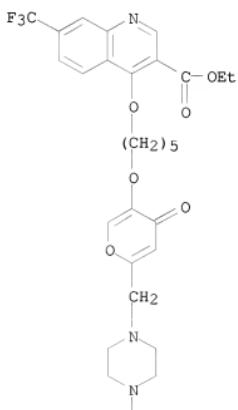
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



RN 754239-86-6 HCPLUS

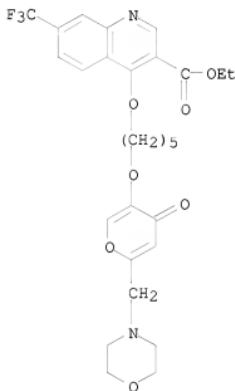
CN 3-Quinolinecarboxylic acid, 4-[(5-[(6-[(4-methyl-1-piperazinyl)methyl]-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)

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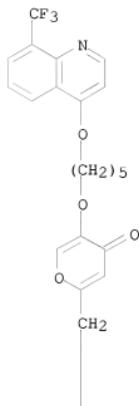




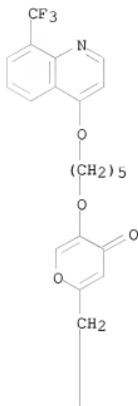
RN 754239-87-7 HCAPLUS  
 CN 3-Quinolincarboxylic acid, 4-[[5-[[6-(4-morpholinylmethyl)-4-oxo-4H-pyran-3-yl]oxy]pentyl]oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)



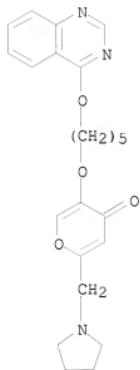
RN 754239-88-8 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[[5-[[8-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



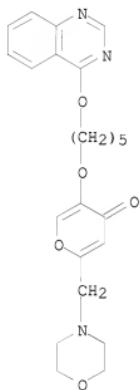
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CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[5-[(8-(trifluoromethyl)-4-quinolinyl)oxy]pentyl]oxy- (CA INDEX NAME)



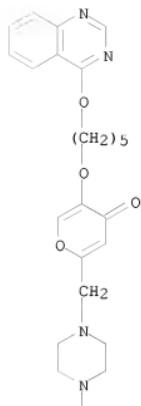
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CN 4H-Pyran-4-one, 2-(1-pyrrolidinylmethyl)-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy- (CA INDEX NAME)



RN 754239-91-3 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(4-morpholinylmethyl)-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)

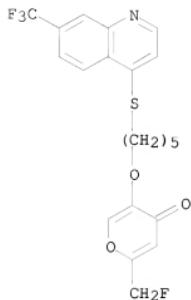


RN 754239-92-4 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)



RN 754239-93-5 HCPLUS

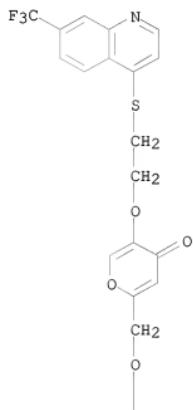
CN 4H-Pyran-4-one, 2-(fluoromethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



RN 754240-07-8 HCPLUS

CN 4H-Pyran-4-one, 2-[(2-oxo-4H-pyran-4-yl)oxy]methyl]-5-[(2-[(7-(trifluoromethyl)-4-quinolinyl]thio)ethoxy]- (CA INDEX NAME)

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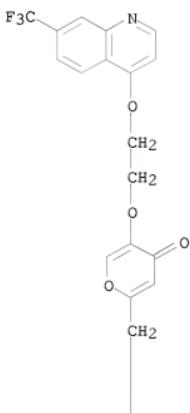
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RN 754240-08-9 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[2-[(7-(trifluoromethyl)-4-quinolinyl)oxy]ethoxy]- (CA INDEX NAME)

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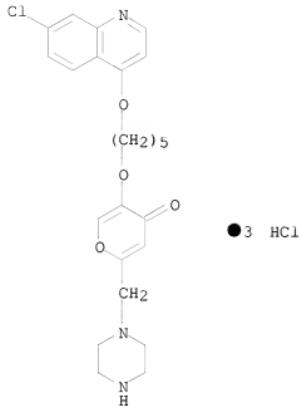


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RN 754240-18-1 HCPLUS

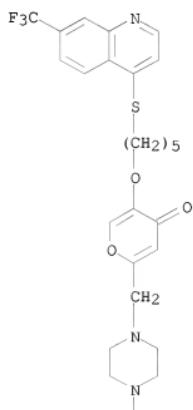
CN 4H-Pyran-4-one, 5-[(5-[(7-chloro-4-quinolinyl)oxy]pentyl)oxy]-2-(1-piperazinylmethyl)-, hydrochloride (1:3) (CA INDEX NAME)



RN 754240-21-6 HCPLUS

CN 4H-Pyran-4-one, 2-[(4-acetyl-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy]- (CA INDEX NAME)

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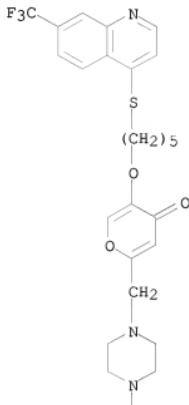
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RN 754240-22-7 HCAPLUS

CN 1-Piperazinecarboxamide, N,N-diethyl-4-[(4-oxo-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]-4H-pyran-2-yl)methyl]- (CA INDEX NAME)

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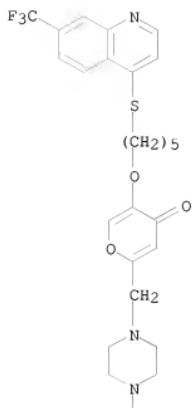


PAGE 2-A



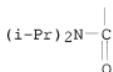
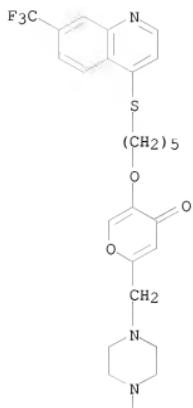
RN 754240-23-8 HCAPLUS

CN 4H-Pyran-4-one, 2-[(4-(2,2-dimethyl-1-oxopropyl)-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



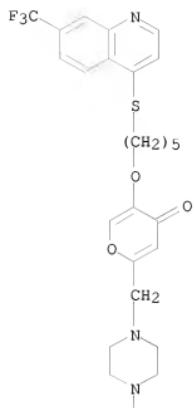
RN 754240-24-9 HCPLUS

CN 1-Piperazinecarboxamide, N,N-bis(1-methylethyl)-4-[(4-oxo-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy)-4H-pyran-2-yl)methyl]-  
(CA INDEX NAME)



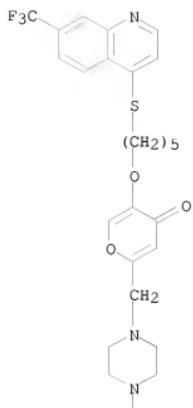
RN 754240-25-0 HCPLUS

CN 4H-Pyran-4-one, 2-[[4-(methylsulfonyl)-1-piperazinyl]methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



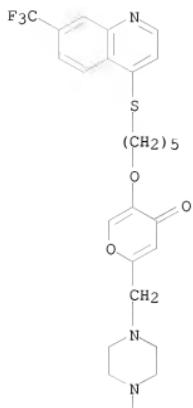
RN 754240-26-1 HCPLUS

CN 1-Piperazinecarboxamide, N-(1,1-dimethylethyl)-4-[(4-oxo-5-[[5-[(7-(trifluoromethyl)-4-quinolinyl)thio]pentyl]oxy]-4H-pyran-2-yl)methyl]-(CA INDEX NAME)



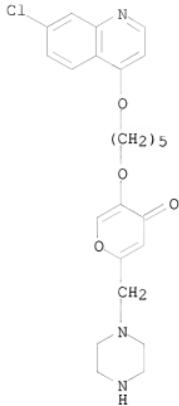
RN 754240-27-2 HCPLUS

CN 1-Piperazinecarboxamide, N-methyl-4-[[4-oxo-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy]-4H-pyran-2-yl]methyl]- (CA INDEX NAME)



RN 754240-28-3 HCAPLUS

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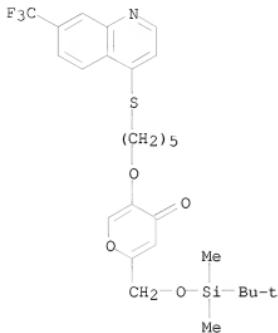


IT 754239-58-2P, 2-[(tert-Butylidimethylsilyloxy)methyl]-5-[(5-[(7-trifluoromethyl)quinolin-4-yl]sulfanyl)pentyl]oxy]-4H-pyran-4-one  
 754239-61-7P, Methanesulfonic acid [4-oxo-5-[(5-[(7-trifluoromethyl)quinolin-4-yl]sulfanyl)pentyl]oxy]-4H-pyran-2-yl]methyl ester 754239-70-8P, 5-[(5-[(6-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-2-(hydroxymethyl)-4H-pyran-4-one] 754239-71-9P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-2-(hydroxymethyl)-4H-pyran-4-one] 754239-72-0P, 4-[(5-[(6-Hydroxymethyl)4-oxo-4H-pyran-3-yl]oxy)pentyl]oxy]-7-trifluoromethylquinoline-3-carboxylic acid ethyl ester 754239-73-1P, 5-[(5-[(8-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-2-(hydroxymethyl)-4H-pyran-4-one] 754239-74-2P, 5-[(5-(Quinazolin-4-yloxy)pentyl]oxy)-2-(hydroxymethyl)-4H-pyran-4-one 754239-75-3P, 5-[(5-[(6-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-76-4P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-77-5P, 4-[(5-[(6-(Methanesulfonyloxy)methyl)-4-oxo-4H-pyran-3-yl]oxy)pentyl]oxy]-7-trifluoromethylquinoline-3-carboxylic acid ethyl ester 754239-78-6P, 5-[(5-[(8-(Trifluoromethyl)quinolin-4-yloxy)pentyl]oxy)-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-79-7P, 5-[(5-(Quinazolin-4-yloxy)pentyl]oxy)-4-oxo-4H-pyran-2-yl)methyl methanesulfonate

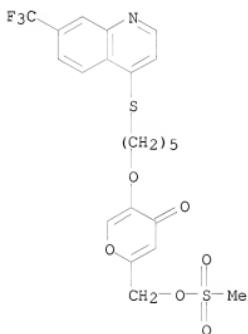
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(intermediate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

BN 754239-58-2 HCAPLUS

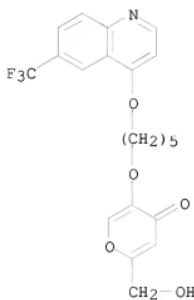
CN 754259-50-2 (CA INDEX NAME)  
 CN 4H-Pyran-4-one, 2-[[[(1,1-dimethylethyl)dimethylsilyl]oxy]methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinylthio]pentyl]oxyl]- (CA INDEX NAME)



RN 754239-61-7 HCAPLUS  
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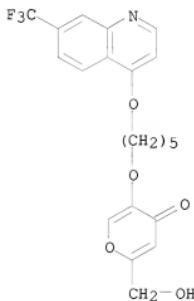


RN 754239-70-8 HCAPLUS  
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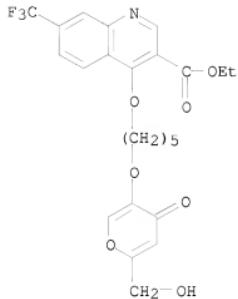
RN 754239-71-9 HCPLUS

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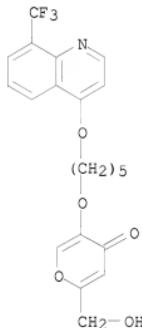
RN 754239-72-0 HCPLUS

CN 3-Quinolinecarboxylic acid, 4-[(5-[(6-(hydroxymethyl)-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)



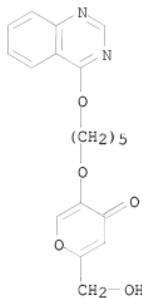
RN 754239-73-1 HCAPLUS

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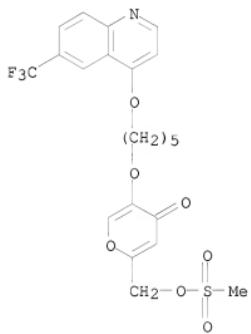
RN 754239-74-2 HCAPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[[5-((4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)



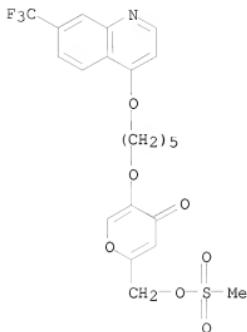
RN 754239-75-3 HCPLUS

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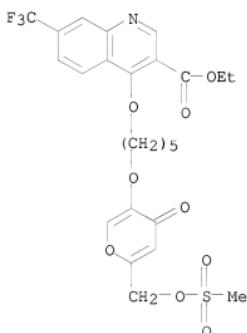


RN 754239-76-4 HCPLUS

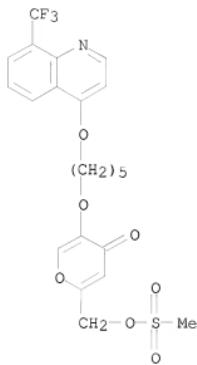
CN 4H-Pyran-4-one, 2-[[5-((methylsulfonyl)oxy)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



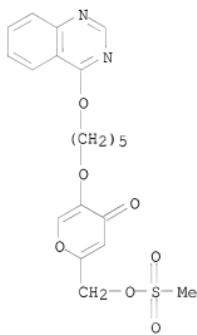
RN 754239-77-5 HCAPLUS  
 CN 3-Quinolinesulfonylic acid, 4-[[5-[[6-[[((methylsulfonyl)oxy)methyl]-4-oxo-4H-pyran-3-yl]oxy]pentyl]oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)



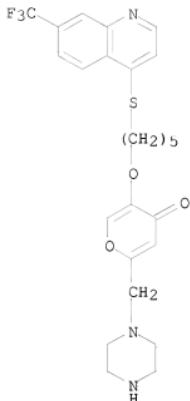
RN 754239-78-6 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[[((methylsulfonyl)oxy)methyl]-5-[[5-[[8-(trifluoromethyl)-4-quinoliny]oxy]pentyl]oxy]- (CA INDEX NAME)



RN 754239-79-7 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(methylsulfonyl)oxy]methyl-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy- (CA INDEX NAME)



IT 754240-20-5, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthio)pentyl]oxy)-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)  
 RN 754240-20-5 HCAPLUS  
 CN 4H-Pyran-4-one, 2-(1-piperazinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinoliny]thio)pentyl]oxy)- (CA INDEX NAME)



L4 ANSWER 5 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1995:608133 HCPLUS

DOCUMENT NUMBER: 123:83751

ORIGINAL REFERENCE NO.: 123:15005a,15008a

TITLE: Mechanism-Based Development of New Antimalarials:  
Synthesis of Derivatives of Artemisinin Attached to  
Iron Chelators

AUTHOR(S): Kamchonwongpaisan, Sumalee; Paitayatat, Sumpan;  
Thebtaranonth, Yodhathai; Wilairat, Prapin; Yuthavong,  
Yongyuth

CORPORATE SOURCE: Faculty of Science, Mahidol University, Bangkok,  
10400, Thailand

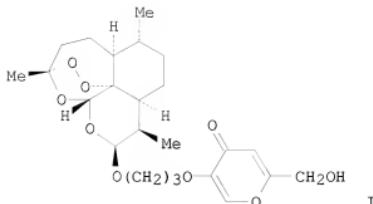
SOURCE: Journal of Medicinal Chemistry (1995), 38(13), 2311-16  
CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

GI



AB Various derivs. of artemisinin covalently linked to iron chelators, e.g. I, were synthesized, and their antimalarial activities were evaluated. Although results show no indication that the presence of an iron chelator in the vicinity of artemisinin potentiates its action, the linked compds. prepared still retain comparable activities to that of artemisinin.

IT 165068-36-0P

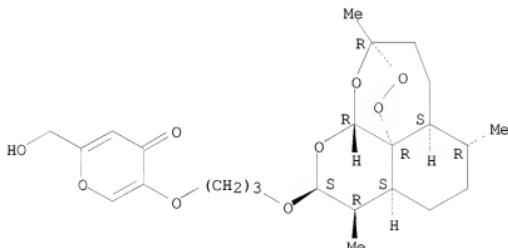
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(mechanism-based development of new antimalarials, synthesis of derivs. of artemisinin attached to iron chelators)

RN 165068-36-0 HCPLUS

CN 4H-Pyran-4-one, 3-[3-[(decahydro-3,6,9-trimethyl-3,12-epoxy-12H-pyran-4,3-yl)-1,2-benzodioxepin-10-yl]oxylpropoxy]-6-(hydroxymethyl)-, [3R-(3 $\alpha$ ,5 $\alpha$ B,6 $\beta$ ,8 $\alpha$ B,9 $\alpha$ ,10 $\alpha$ ,12 $\beta$ ,12aR\*)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



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10541328

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DICTIONARY FILE UPDATES: 2 AUG 2008 HIGHEST RN 1037774-47-2

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Uploading C:\Program Files\Stnexp\Queries\10541328a.str



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ring nodes :  
1 2 3 4 5 6  
chain bonds :  
3-12 4-7 9-10 9-12 10-15

10541328

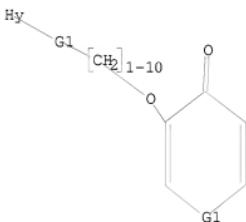
ring bonds :  
1-2 1-6 2-3 3-4 4-5 5-6  
exact/norm bonds :  
1-2 1-6 2-3 3-4 3-12 4-5 4-7 5-6 9-10 9-12 10-15  
isolated ring systems :  
containing 1 :

G1:O,S,N

Match level :  
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 9:CLASS 10:Atom 12:CLASS  
15:Atom

L6 STRUCTURE UPLOADED

=> d 16  
L6 HAS NO ANSWERS  
L6 STR



G1 O,S,N

Structure attributes must be viewed using STN Express query preparation.

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SAMPLE SCREEN SEARCH COMPLETED - 28332 TO ITERATE

7.1% PROCESSED 2000 ITERATIONS 0 ANSWERS  
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 556571 TO 576709  
PROJECTED ANSWERS: 0 TO 0

L7 0 SEA SSS SAM L6

10541328

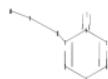
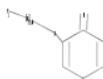
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FULL SCREEN SEARCH COMPLETED - 575864 TO ITERATE

100.0% PROCESSED 575864 ITERATIONS  
SEARCH TIME: 00.00.08

85 ANSWERS

L8 85 SEA SSS FUL L6

=>  
Uploading C:\Program Files\Stnexp\Queries\10541328b.str



chain nodes :

7 9 10 12

ring nodes :

1 2 3 4 5 6

chain bonds :

3-12 4-7 9-10 9-12

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

1-2 1-6 2-3 3-4 3-12 4-5 4-7 5-6 9-10 9-12

isolated ring systems :

containing 1 :

G1:O,S,N

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 9:CLASS 10:Atom 12:CLASS

L9 STRUCTURE UPLOADED

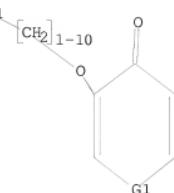
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L9 HAS NO ANSWERS

L9

STR

G1



G1

G1 O,S,N

Structure attributes must be viewed using STN Express query preparation.

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 SAMPLE SCREEN SEARCH COMPLETED - 28332 TO ITERATE

7.1% PROCESSED 2000 ITERATIONS 1 ANSWERS  
 INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)  
 SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
 BATCH \*\*COMPLETE\*\*  
 PROJECTED ITERATIONS: 556571 TO 576709  
 PROJECTED ANSWERS: 58 TO 508

L10 1 SEA SSS SAM L9

=> s 19 sss full  
 FULL SEARCH INITIATED 10:47:55 FILE 'REGISTRY'  
 FULL SCREEN SEARCH COMPLETED - 575864 TO ITERATE

100.0% PROCESSED 575864 ITERATIONS 218 ANSWERS  
 SEARCH TIME: 00.00.04

L11 218 SEA SSS FUL L9

=> FIL HCAPLUS			
COST IN U.S. DOLLARS	SINCE FILE	TOTAL	
	ENTRY	SESSION	
FULL ESTIMATED COST	357.18	581.83	
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	ENTRY	SESSION	
CA SUBSCRIBER PRICE	0.00	-4.00	

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FILE COVERS 1907 - 4 Aug 2008 VOL 149 ISS 6  
FILE LAST UPDATED: 3 Aug 2008 (20080803/ED)

HCPlus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2008.

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L2                   0 S L1  
L3                   69 S L1 SSS FULL

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L4                   5 S L3  
L5                   1 S L4 AND PY<=2003

FILE 'REGISTRY' ENTERED AT 10:46:19 ON 04 AUG 2008

L6                   STRUCTURE UPLOADED  
L7                   0 S L6  
L8                   85 S L6 SSS FULL  
L9                   STRUCTURE UPLOADED  
L10                  1 S L9  
L11                  218 S L9 SSS FULL

FILE 'HCAPLUS' ENTERED AT 10:48:03 ON 04 AUG 2008

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=> s 111  
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=> s 112 and py<=2003  
24005635 PY<=2003  
L14                  1 L12 AND PY<=2003

=> s 113 and py<=2003  
24005635 PY<=2003

L15 17 L13 AND PY&lt;=2003

=&gt; s l15 and p/dt

6298780 P/DT

L16 11 L15 AND P/DT

=&gt; s l16 and us/pc

1822860 US/PC

L17 5 L16 AND US/PC

=&gt; d l12 ibib abs hitstr tot

L12 ANSWER 1 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2008:622524 HCPLUS

TITLE: Characterization of EHT 1864, a novel small molecule inhibitor of Rac family small GTPases  
Onesto, Cercina; Shutes, Adam; Picard, Virginie; Schweihofer, Fabien; Der, Channing J.

CORPORATE SOURCE: Lineberger Comprehensive Cancer Center, Department of Pharmacology, University of North Carolina at Chapel Hill, Chapel Hill, NC, USA

SOURCE: Methods in Enzymology (2008), 439(Small GTPases in Disease, Part B), 111-129  
CODEN: MENZAU; ISSN: 0076-6879

PUBLISHER: Elsevier

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

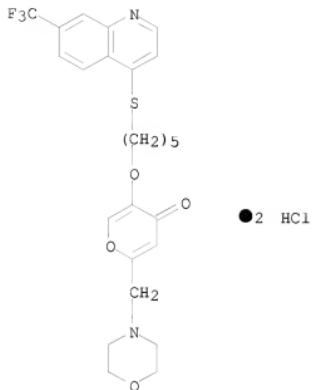
AB A review. There is now considerable exptl. evidence that aberrant activation of Rho family small GTPases promotes uncontrolled proliferation, invasion, and metastatic properties of human cancer cells. Therefore, there is considerable interest in the development of small mol. inhibitors of Rho GTPase function. However, to date, most efforts have focused on inhibitors that block Rho GTPase function indirectly, either by targeting enzymes involved in post-translational processing or downstream protein kinase effectors. We have reported the identification and characterization of the EHT 1864 small mol. as an inhibitor of Rac family small GTPases, placing Rac1 in an inert and inactive state and then impairing Rac1-mediated functions *in vivo*. Our work suggests that EHT 1864 selectively inhibits Rac1 downstream signaling and cellular transformation by a novel mechanism involving guanine nucleotide displacement. This chapter provides the details for some of the biochem. and biol. methods used to characterize the mode of action of EHT 1864 on Rac1 and its impact on Rac1-dependent cellular functions.

IT 754240-09-0, EHT1864

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(biochem. and biol. methods may be useful to characterize Rho GTPase specificity and mechanism of action of EHT 1864 on Rac1 and its impact on Rac1-dependent cellular function in mouse)

RN 754240-09-0 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:2) (CA INDEX NAME)



L12 ANSWER 2 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 2007:1368320 HCPLUS  
 DOCUMENT NUMBER: 148:232138  
 TITLE: Specificity and Mechanism of Action of EHT 1864, a Novel Small Molecule Inhibitor of Rac Family Small GTPases  
 AUTHOR(S): Shutes, Adam; Onesto, Cercina; Picard, Virginie; Leblond, Bertrand; Schweighoffer, Fabien; Der, Channing J.  
 CORPORATE SOURCE: Lineberger Comprehensive Cancer Center, University of North Carolina, Chapel Hill, NC, 27599, USA  
 SOURCE: Journal of Biological Chemistry (2007), 282(49), 35666-35678  
 CODEN: JBCHA3; ISSN: 0021-9258  
 PUBLISHER: American Society for Biochemistry and Molecular Biology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB There is now considerable exptl. evidence that aberrant activation of Rho family small GTPases promotes the uncontrolled proliferation, invasion, and metastatic properties of human cancer cells. Therefore, there is considerable interest in the development of small mol. inhibitors of Rho GTPase function. However, to date, most efforts have focused on inhibitors that indirectly block Rho GTPase function, by targeting either enzymes involved in post-translational processing or downstream protein kinase effectors. We recently determined that the EHT 1864 small mol. can inhibit Rac function *in vivo*. In this study, we evaluated the biol. and biochem. specificities and biochem. mechanism of action of EHT 1864. We determined that EHT 1864 specifically inhibited Rac1-dependent platelet-derived growth factor-induced lamellipodia formation. Furthermore, our biochem. analyses with recombinant Rac proteins found that EHT 1864 possesses high

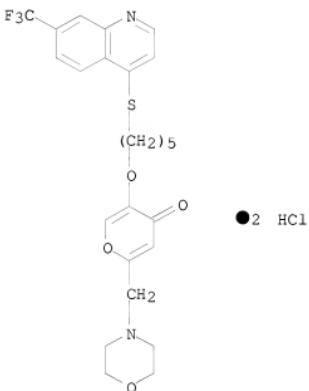
affinity binding to Rac1, as well as the related Rac1b, Rac2, and Rac3 isoforms, and this association promoted the loss of bound nucleotide, inhibiting both guanine nucleotide association and Tiam1 Rac guanine nucleotide exchange factor-stimulated exchange factor activity *in vitro*. EHT 1864 therefore places Rac in an inert and inactive state, preventing its engagement with downstream effectors. Finally, we evaluated the ability of EHT 1864 to block Rac-dependent growth transformation, and we determined that EHT 1864 potently blocked transformation caused by constitutively activated Rac1, as well as Rac-dependent transformation caused by Tiam1 or Ras. Taken together, our results suggest that EHT 1864 selectively inhibits Rac downstream signaling and transformation by a novel mechanism involving guanine nucleotide displacement.

IT 754240-09-0, EHT 1864

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(EHT 1864 specifically inhibits Rac1-dependent platelet-derived growth factor-induced lamellipodia formation)

RN 754240-09-0 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy)-, hydrochloride (1:2) (CA INDEX NAME)



REFERENCE COUNT: 61 THERE ARE 61 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2007:525951 HCPLUS

DOCUMENT NUMBER: 147:143245

TITLE: Chemotherapy of leishmaniasis. Part V: Synthesis and *in vitro* bioevaluation of novel pyridinone derivatives

AUTHOR(S): Pandey, Susmita; Suryawanshi, S. N.; Nishi; Goyal, Neena; Gupta, Suman

CORPORATE SOURCE: Division of Medicinal Chemistry, Central Drug Research Institute, Lucknow, Uttar Pradesh, 226001, India

SOURCE: European Journal of Medicinal Chemistry (2007), 42(5), 669-674

CODEN: EJMC5A; ISSN: 0223-5234

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 147:143245

AB Alkylation of 3-hydroxy-2-methyl-4H-pyran-4-one with 1,5-dibromopentane gave dimeric  $\gamma$ -pyrone (2), which was recyclized with substituted anilines or cyclohexylamine to yield 1,1'-diaryl-2,2'-dimethyl-3,3'-pentamethylenedioxy-4,4'-bipyridinones (3a,b, d-j; aryl = 2,6-Me<sub>2</sub>C<sub>6</sub>H<sub>3</sub>, 2,3-Me<sub>2</sub>C<sub>6</sub>H<sub>3</sub>, 4-MeC<sub>6</sub>H<sub>4</sub>, Ph, PhCH<sub>2</sub>, 4-FC<sub>6</sub>H<sub>4</sub>, 4-ClC<sub>6</sub>H<sub>4</sub>, 4-NO<sub>2</sub>C<sub>6</sub>H<sub>4</sub>, 2-MeOC<sub>6</sub>H<sub>4</sub>) and 1,1'-dicyclohexyl derivative (3c), together with the corresponding pyridinone-pyrone derivs., 1-Ar-2-methyl-3-[5-(2-methyl-4-oxo-4H-pyran-3-yloxy)pentyl]oxy]-4(4H)-pyridinones (4a-j, same Ar). The novel 2-substituted pyridinone derivs. were screened towards *in vitro* for their activity against leishmania antipromastigote and antiamastigote activity profile, exhibiting good activity in some cases (3a, 3b; aryl = 2,6-Me<sub>2</sub>C<sub>6</sub>H<sub>3</sub>, 2,3-Me<sub>2</sub>C<sub>6</sub>H<sub>3</sub>, 4i, 4j, Ar = 4-NO<sub>2</sub>C<sub>6</sub>H<sub>4</sub>, 2-MeOC<sub>6</sub>H<sub>4</sub>).

IT 943737-55-1P 943737-57-3P 943737-59-5P

943737-61-9P 943737-63-1P 943737-64-2P

943737-66-4P 943737-68-6P 943737-70-0P

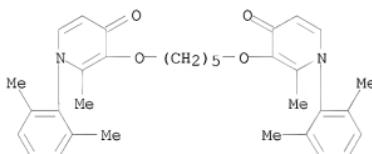
943737-72-2P 943737-74-4P 943737-76-6P

943737-77-7P 943737-79-9P

RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(antileishmaniasis activity; preparation of pentamethylenedioxy-bridged bis-4-pyridinones and pyridinone-pyrone as antimicrobial agents against leishmaniasis)

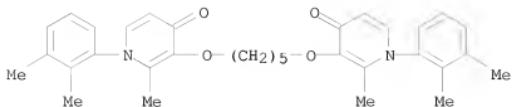
RN 943737-55-1 HCPLUS

CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[1-(2,6-dimethylphenyl)-2-methyl- (CA INDEX NAME)

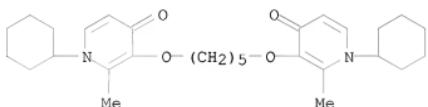


RN 943737-57-3 HCPLUS

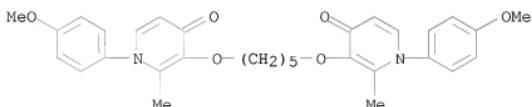
CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[1-(2,3-dimethylphenyl)-2-methyl- (CA INDEX NAME)



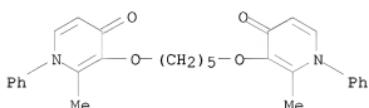
RN 943737-59-5 HCAPLUS  
 CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[1-cyclohexyl-2-methyl-  
 (CA INDEX NAME)



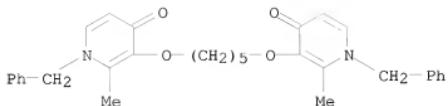
RN 943737-61-9 HCAPLUS  
 CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[1-(4-methoxyphenyl)-2-methyl-  
 (CA INDEX NAME)



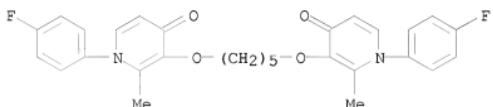
RN 943737-63-1 HCAPLUS  
 CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[2-methyl-1-phenyl-  
 (CA INDEX NAME)



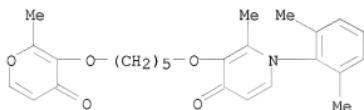
RN 943737-64-2 HCAPLUS  
 CN 4(1H)-Pyridinone, 3,3'-[1,5-pentanediylbis(oxy)]bis[2-methyl-1-(phenylmethyl)-  
 (CA INDEX NAME)



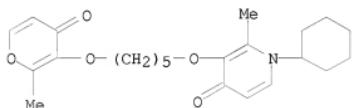
RN 943737-66-4 HCAPLUS  
 CN 4(1H)-Pyridinone, 3,3'-(1,5-pentanediylbis(oxy))bis[1-(4-fluorophenyl)-2-methyl- (CA INDEX NAME)



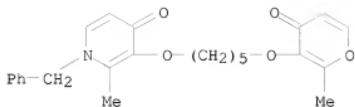
RN 943737-68-6 HCAPLUS  
 CN 4(1H)-Pyridinone, 1-(2,6-dimethylphenyl)-2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]- (CA INDEX NAME)



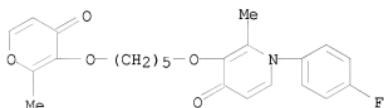
RN 943737-70-0 HCAPLUS  
 CN 4(1H)-Pyridinone, 1-cyclohexyl-2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]- (CA INDEX NAME)



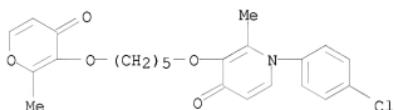
RN 943737-72-2 HCAPLUS  
 CN 4(1H)-Pyridinone, 2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-1-(phenylmethyl)- (CA INDEX NAME)



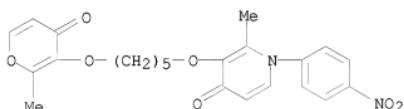
RN 943737-74-4 HCAPLUS  
 CN 4(1H)-Pyridinone, 1-(4-fluorophenyl)-2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyloxy)- (CA INDEX NAME)



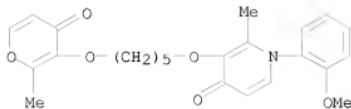
RN 943737-76-6 HCAPLUS  
 CN 4(1H)-Pyridinone, 1-(4-chlorophenyl)-2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyloxy)- (CA INDEX NAME)



RN 943737-77-7 HCAPLUS  
 CN 4(1H)-Pyridinone, 2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyloxy)-1-(4-nitrophenyl)- (CA INDEX NAME)



RN 943737-79-9 HCAPLUS  
 CN 4(1H)-Pyridinone, 1-(2-methoxyphenyl)-2-methyl-3-[(5-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]pentyloxy)- (CA INDEX NAME)

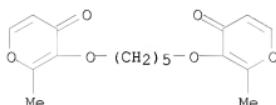


IT 943737-53-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
 (preparation of pentamethylenedioxy-bridged bis-4-pyridinones and pyridinone-pyrones as antimicrobial agents against leishmaniasis)

RN 943737-53-9 HCPLUS

CN 4H-Pyran-4-one, 3,3'-[1,5-pentanediyloxy]bis[2-methyl- (CA INDEX NAME)



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 4 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:1191062 HCPLUS

DOCUMENT NUMBER: 144:68139

TITLE: RAC1 Inhibition Targets Amyloid Precursor Protein Processing by  $\gamma$ -Secretase and Decreases A $\beta$  Production in Vitro and in Vivo

AUTHOR(S): Desire, Laurent; Bourdin, Jerome; Loiseau, Nadia; Peillon, Helene; Picard, Virginie; De Oliveira, Catherine; Bachelot, Florence; Leblond, Bertrand; Taverne, Thierry; Beausoleil, Eric; Lacome, Sandrine; Drouin, Dominique; Schweighoffer, Fabien

CORPORATE SOURCE: Exonhit Therapeutics, Paris, 75013, Fr.

SOURCE: Journal of Biological Chemistry (2005), 280(45), 37516-37525

CODEN: JBCHA3; ISSN: 0021-9258

PUBLISHER: American Society for Biochemistry and Molecular Biology

DOCUMENT TYPE: Journal  
LANGUAGE: English

AB  $\beta$ -Amyloid peptides (A $\beta$ ) that form the senile plaques of Alzheimer disease consist mainly of 40- and 42-amino acid (A $\beta$  40 and A $\beta$  42) peptides generated from the cleavage of the amyloid precursor protein (APP). Generation of A $\beta$  involves  $\beta$ -secretase and  $\gamma$ -secretase activities and is regulated by membrane trafficking of the proteins involved in A $\beta$  production. Here we describe a new small mol., EHT 1864, which blocks the RAC1 signaling pathways. In vitro, EHT 1864 blocks A $\beta$  40 and A $\beta$  42 production but does not impact

SAPP $\alpha$  levels and does not inhibit  $\beta$ -secretase. Rather, EHT 1864 modulates APP processing at the level of  $\gamma$ -secretase to prevent A $\beta$  40 and A $\beta$  42 generation. This effect does not result from a direct inhibition of the  $\gamma$ -secretase activity and is specific for APP cleavage, since EHT 1864 does not affect Notch cleavage. In vivo, EHT 1864 significantly reduces A $\beta$  40 and A $\beta$  42 levels in guinea pig brains at a threshold that is compatible with delaying plaque accumulation and/or clearing the existing plaque in brain. EHT 1864 is the first derivative of a new chemical series that consists of candidates for inhibiting A $\beta$  formation in the brain of AD patients. Our findings represent the first pharmacol. validation of Rac1 signaling as a target for developing novel therapies for Alzheimer disease.

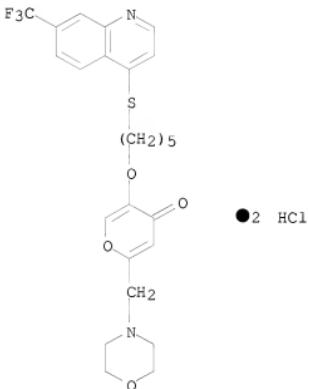
IT 754240-09-0

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(EHT 1864; EHT 1864 blocked A $\beta$ 1-40 and A $\beta$ 1-42 production)

RN 754240-09-0 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thiopentyl)oxy]-, hydrochloride (1:2) (CA INDEX NAME)



REFERENCE COUNT: 55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 5 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:740320 HCPLUS

DOCUMENT NUMBER: 141:260557

TITLE: Preparation of novel antiproliferative and

antiangiogenic agents, in particular  
quinoline-derivatized pyranones, for treating cell  
proliferative diseases, retinopathies and arthritis  
Leblond, Bertrand; Petit, Silvere; Picard, Virginie;  
Taverne, Thierry; Schweighoffer, Fabien

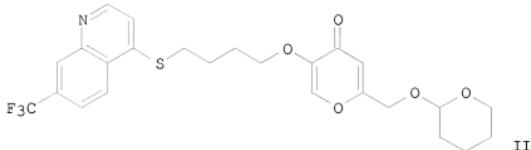
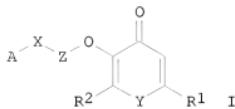
INVENTOR(S):

PATENT ASSIGNEE(S): Exonhit Therapeutics Sa, Fr.  
 SOURCE: PCT Int. Appl., 156 pp.

DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004076445	A2	20040910	WO 2004-IB926	20040227
WO 2004076445	A3	20050106		
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EP 1471063	A1	20041027	EP 2003-290490	20030228
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AU 2004215577	A1	20040910	AU 2004-215577	20040227
CA 2516239	A1	20040910	CA 2004-2516239	20040227
EP 1597253	A2	20051123	EP 2004-715422	20040227
EP 1597253	B1	20060809		
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CN 1747952	A	20060315	CN 2004-80003820	20040227
JP 2006519221	T	20060824	JP 2006-502497	20040227
AT 335734	T	20060915	AT 2004-715422	20040227
US 20060183749	A1	20060817	US 2005-541328	20050830
PRIORITY APPLN. INFO.:			EP 2003-290490	A 20030228
			WO 2004-IB926	W 20040227

OTHER SOURCE(S): MARPAT 141:260557  
 GI



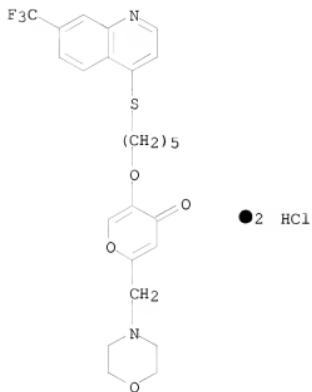
AB Title compds. I [wherein R1 = [(tetrahydropyran-2-yl)oxy]methyl, CH2-B, (morpholin-4-yl)methyl, pyrrolidin-1-ylmethyl, etc.; B = halo, OH, OCH2OMe, OCH2OCH2CH2OMe, OS(=O)(=O)alkyl, OTBDMS; R2 = H, alk(en)yl; X, Y = independently O, S, NH and derivs.; A = quinolin-4-yl, quinolin-8-yl, benzo[b]thiophen-7-yl, quinazolin-4-yl; Z = (CH2)*n*, optionally interrupted by a heteroatom, C(=O) or arylidialkyl, especially xyleneyl, group; *n* = 1-10; their tautomers, optical and geometrical isomers, racemates, salts, hydrates and mixts.] were prepared as antiproliferative agents and angiogenesis inhibitors. Nine biol. assays are given. For example, II was prepared, in 2 steps, from pyranone III, 1,4-dibromobutane, and 7-(trifluoromethyl)-4-quinolinethiol. In an *in vitro* cell viability assay, selected I showed an IC50 < 4  $\mu$ M and < 9  $\mu$ M against HCT116 and MDA-MB-231 tumoral cell lines, demonstrating their cytostatic mode of action. I are useful for treating various diseases associated with abnormal cell proliferation, including cancer, especially leukemia, or associated with unregulated angiogenesis including growth and metastasis of solid tumors, ocular diseases, especially retinopathies, or arthritis.

IT 754240-09-0P, 5-[(5-[7-(Trifluoromethyl)quinolin-4-ylthioliopentyl]oxy)-2-(morpholinomethyl)-4H-pyran-4-one dihydrochloride  
 754240-17-0P, 5-[(5-[7-(Trifluoromethyl)quinolin-4-ylthioliopentyl]oxy)-2-[(4-methylpiperazin-1-yl)methyl]-4H-pyran-4-one trihydrochloride 754240-19-2P, 5-[(5-[7-(Trifluoromethyl)quinolin-4-ylthioliopentyl]oxy)-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one trihydrochloride

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

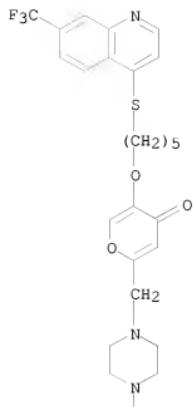
RN 754240-09-0 HCPLUS  
 CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[7-(trifluoromethyl)-4-

quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:2) (CA INDEX NAME)



RN 754240-17-0 HCPLUS

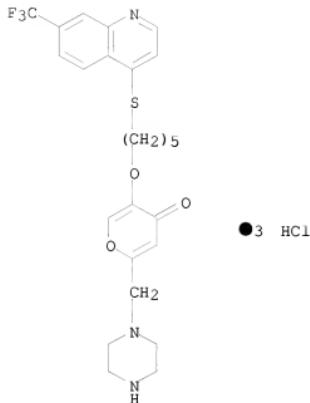
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]-, hydrochloride (1:3) (CA INDEX NAME)



● 3 HCl

RN 754240-19-2 HCPLUS

CN 4H-Pyran-4-one, 2-(1-piperazinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]-, hydrochloride (1:3) (CA INDEX NAME)



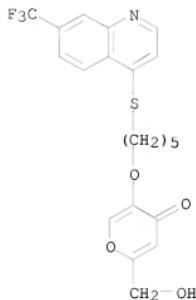
IT 754239-59-3P, 2-Hydroxymethyl-5-[(5-[7-(trifluoromethyl)quinolin-4-ylsulfanyl]pentyl)oxy]-4H-pyran-4-one 754240-15-8P, tert-Butyl 4-[(5-[5-[7-(trifluoromethyl)quinolin-4-ylthio]pentyl)oxy]-4-oxo-4H-pyran-2-ylmethyl]piperazine-1-carboxylate 754240-16-9P, tert-Butyl 4-[(5-[5-(7-chloroquinolin-4-yl)pentyl)oxy]-4-oxo-4H-pyran-2-ylmethyl]piperazine-1-carboxylate

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

RN 754239-59-3 HCAPLUS

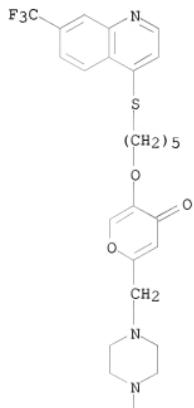
CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[7-(trifluoromethyl)-4-quinolinyl]thiopentyl)oxy]- (CA INDEX NAME)



RN 754240-15-8 HCPLUS

CN 1-Piperazinecarboxylic acid, 4-[(4-oxo-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl)thiolpentyl]oxy)-4H-pyran-2-yl]methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

PAGE 1-A



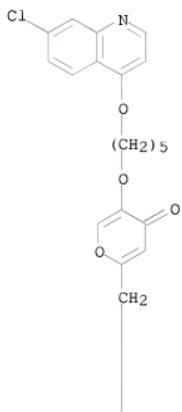
PAGE 2-A



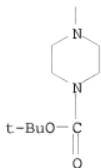
RN 754240-16-9 HCPLUS

CN 1-Piperazinecarboxylic acid, 4-[[5-[[5-[(7-chloro-4-quinolinyl)oxypentyl]oxy]-4-oxo-4H-pyran-2-yl]methyl]-, 1,1-dimethylethyl ester (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



IT 754239-47-9P, 5-[[5-((6-Fluoro-2-methylquinolin-4-yloxy)pentyl)oxy]-2-[(tetrahydropyran-2-yl)oxy]methyl]-4H-pyran-4-one 754239-48-0P



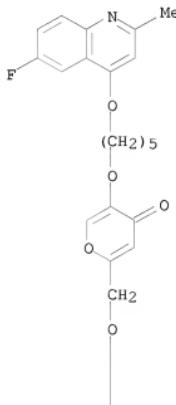
(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy]-2-[(4-acetylpirazin-1-yl)methyl]-4H-pyran-4-one 754240-22-7P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N,N-diethylpiperazine-1-carboxamide 754240-23-8P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-2-[(4-(pivaloyl)piperazin-1-yl)methyl]-4H-pyran-4-one 754240-24-9P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N,N-diisopropylpiperazine-1-carboxamide 754240-25-0P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-2-[(4-Methylsulfonylpiperazin-1-yl)methyl]-4H-pyran-4-one 754240-26-1P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N-tert-butylpiperazine-1-carboxamide 754240-27-2P, 4-[(5-[(5-[(7-(Trifluoromethyl)quinolin-4-ylthiopentyl]oxy)-4-oxo-4H-pyran-2-yl]methyl)-N-methylpiperazine-1-carboxamide 754240-28-3P, 5-[(5-[(7-Chloroquinolin-4-yloxy)pentyl]oxy)-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

RN 754239-47-9 HCAPLUS

CN 4H-Pyran-4-one, 5-[(5-[(6-fluoro-2-methyl-4-quinolinyl)oxy]pentyl]oxy)-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl]- (CA INDEX NAME)

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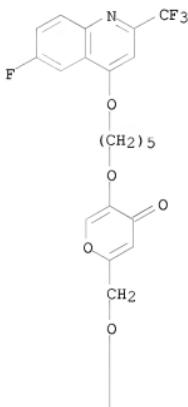


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RN 754239-48-0 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[[5-[(6-fluoro-2-(trifluoromethyl)-4-quinolinyl)oxy]pentyl]oxy]-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl- (CA INDEX NAME)

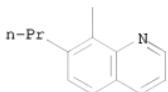
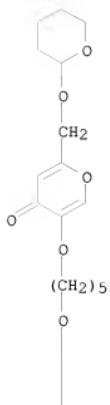
PAGE 1-A



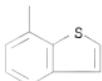
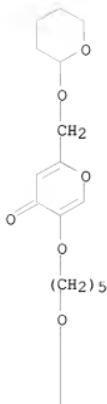
PAGE 2-A



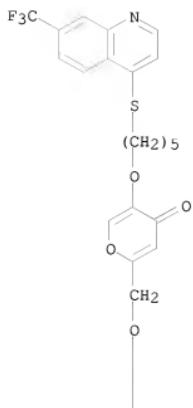
RN 754239-49-1 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[[5-[(7-propyl-8-quinolinyl)oxy]pentyl]oxy]-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl- (CA INDEX NAME)



RN 754239-50-4 HCPLUS  
CN 4H-Pyran-4-one, 5-[[5-(benzo[b]thien-7-yl)oxy]pentyl]oxy]-2-[[[(tetrahydro-2H-pyran-2-yl)oxy]methyl]- (CA INDEX NAME)

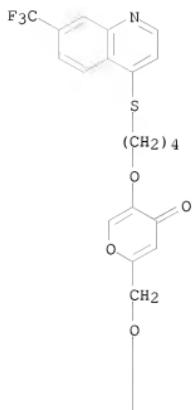


RN 754239-51-5 HCPLUS  
CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl)thio]pentyl)oxy]- (CA INDEX NAME)



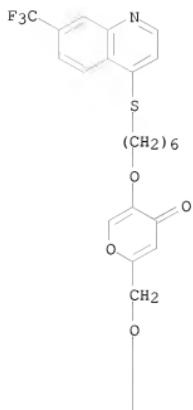
RN 754239-53-7 HCPLUS

CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[4-[(7-(trifluoromethyl)-4-quinolinyl]thio]butoxy]- (CA INDEX NAME)



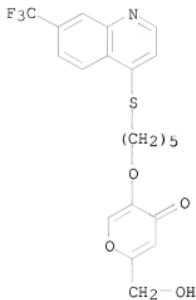
RN 754239-55-9 HCPLUS

CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(6-[(7-(trifluoromethyl)-4-quinolinyl]thio)hexyl]oxy]- (CA INDEX NAME)



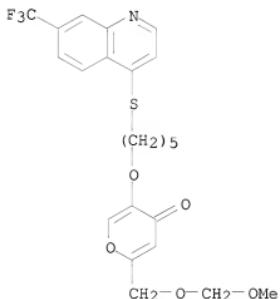
RN 754239-56-0 HCPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy]-, hydrochloride (1:1) (CA INDEX NAME)

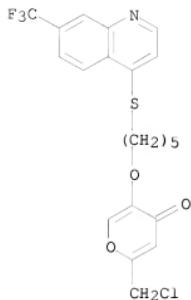


● HCl1

RN 754239-60-6 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(methoxymethoxy)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



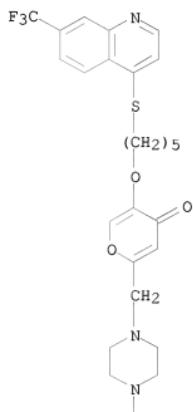
RN 754239-62-8 HCAPLUS  
 CN 4H-Pyran-4-one, 2-(chloromethyl)-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



RN 754239-63-9 HCPLUS

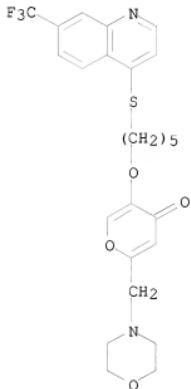
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy]methyl] (CA INDEX NAME)

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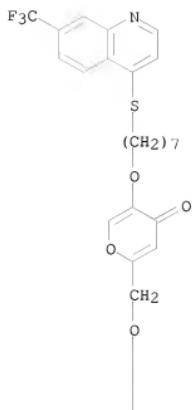




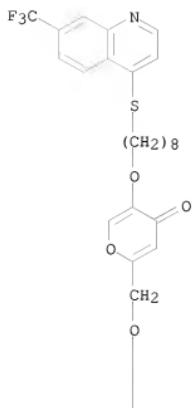
RN 754239-64-0 HCAPLUS  
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinoliny]thio)pentyl]oxy]- (CA INDEX NAME)



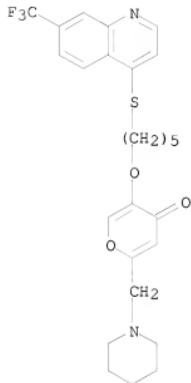
RN 754239-66-2 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(7-[(7-(trifluoromethyl)-4-quinoliny]thio)heptyl]oxy]- (CA INDEX NAME)



RN 754239-68-4 HCPLUS  
CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl]-5-[(8-[(7-(trifluoromethyl)-4-quinoliny]thio]octyl]oxy]- (CA INDEX NAME)

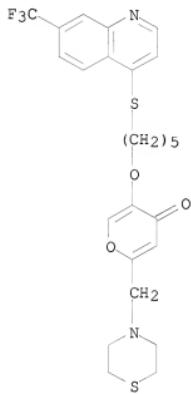


RN 754239-80-0 HCPLUS  
CN 4H-Pyran-4-one, 2-(1-piperidinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thiopentyl]oxy]- (CA INDEX NAME)



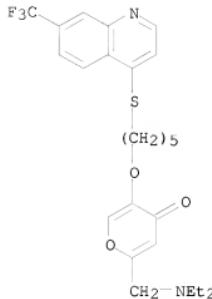
RN 754239-81-1 HCPLUS

CN 4H-Pyran-4-one, 2-(4-thiomorpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



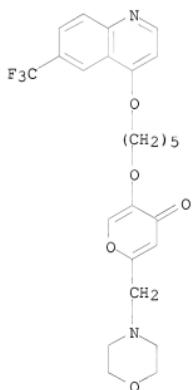
RN 754239-82-2 HCPLUS

CN 4H-Pyran-4-one, 2-[(diethylamino)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



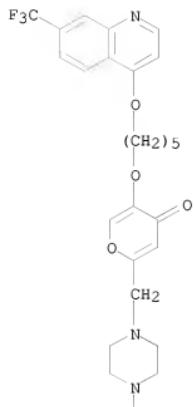
RN 754239-83-3 HCAPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(6-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



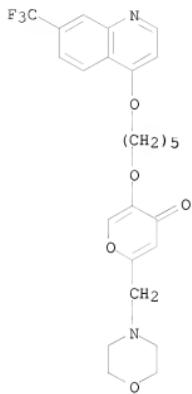
RN 754239-84-4 HCAPLUS

CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



RN 754239-85-5 HCPLUS

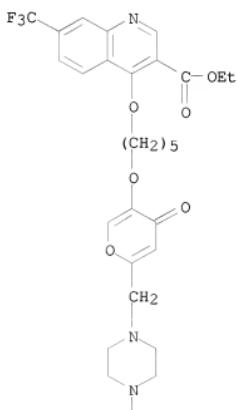
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



RN 754239-86-6 HCPLUS

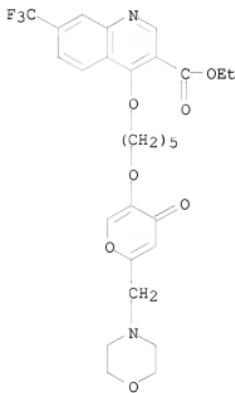
CN 3-Quinolinecarboxylic acid, 4-[(5-[(6-[(4-methyl-1-piperazinyl)methyl]-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)

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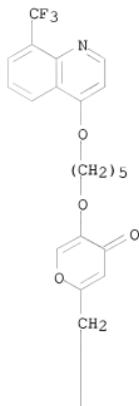




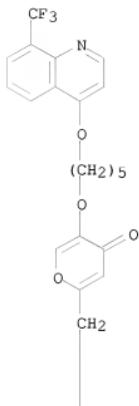
RN 754239-87-7 HCAPLUS  
 CN 3-Quinolincarboxylic acid, 4-[[5-[[6-(4-morpholinylmethyl)-4-oxo-4H-pyran-3-yl]oxy]pentyl]oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)



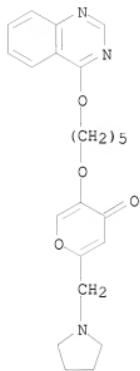
RN 754239-88-8 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[[5-[[8-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



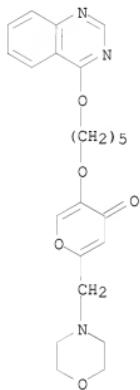
RN 754239-89-9 HCPLUS  
CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[5-[(8-(trifluoromethyl)-4-quinolinyl)oxy]pentyl]oxy- (CA INDEX NAME)



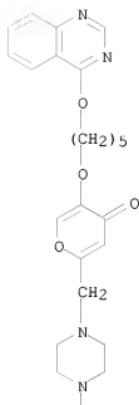
RN 754239-90-2 HCPLUS  
CN 4H-Pyran-4-one, 2-(1-pyrrolidinylmethyl)-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy- (CA INDEX NAME)



RN 754239-91-3 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(4-morpholinylmethyl)-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)

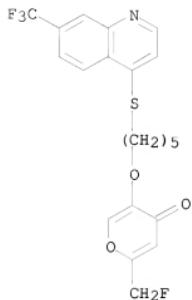


RN 754239-92-4 HCAPLUS  
CN 4H-Pyran-4-one, 2-[(4-methyl-1-piperazinyl)methyl]-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)



RN 754239-93-5 HCPLUS

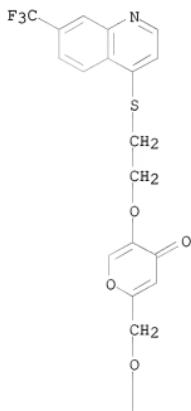
CN 4H-Pyran-4-one, 2-(fluoromethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



RN 754240-07-8 HCPLUS

CN 4H-Pyran-4-one, 2-[([(tetrahydro-2H-pyran-2-yl)oxy]methyl)-5-[(2-[(7-(trifluoromethyl)-4-quinolinyl]thio)ethoxy]- (CA INDEX NAME)

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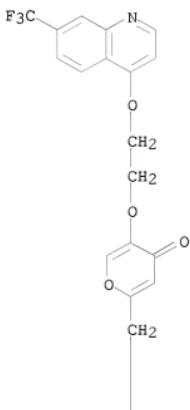
PAGE 2-A



RN 754240-08-9 HCPLUS

CN 4H-Pyran-4-one, 2-(4-morpholinylmethyl)-5-[2-[(7-(trifluoromethyl)-4-quinolinyl)oxy]ethoxy]- (CA INDEX NAME)

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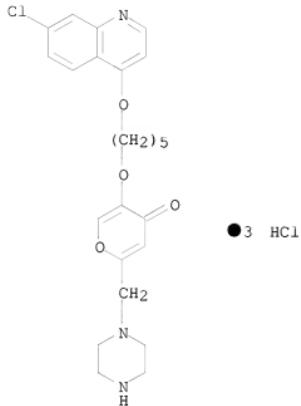


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RN 754240-18-1 HCPLUS

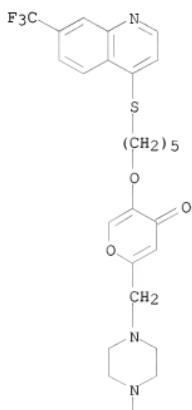
CN 4H-Pyran-4-one, 5-[(5-[(7-chloro-4-quinolinyl)oxy]pentyl)oxy]-2-(1-piperazinylmethyl)-, hydrochloride (1:3) (CA INDEX NAME)



RN 754240-21-6 HCPLUS

CN 4H-Pyran-4-one, 2-[(4-acetyl-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy]- (CA INDEX NAME)

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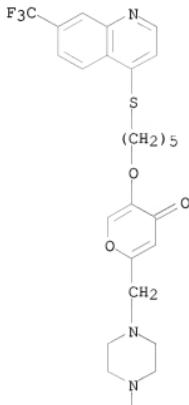
PAGE 2-A



RN 754240-22-7 HCAPLUS

CN 1-Piperazinecarboxamide, N,N-diethyl-4-[(4-oxo-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]-4H-pyran-2-yl)methyl]- (CA INDEX NAME)

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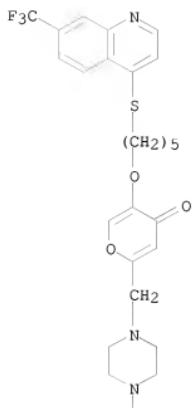


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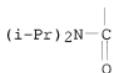
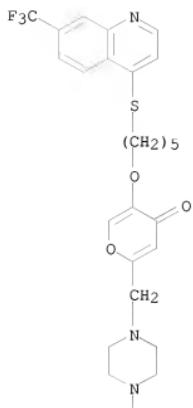
RN 754240-23-8 HCAPLUS

CN 4H-Pyran-4-one, 2-[(4-(2,2-dimethyl-1-oxopropyl)-1-piperazinyl)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio]pentyl)oxy]- (CA INDEX NAME)



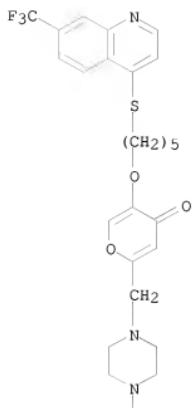
RN 754240-24-9 HCPLUS

CN 1-Piperazinecarboxamide, N,N-bis(1-methylethyl)-4-[(4-oxo-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy)-4H-pyran-2-yl)methyl]-(CA INDEX NAME)



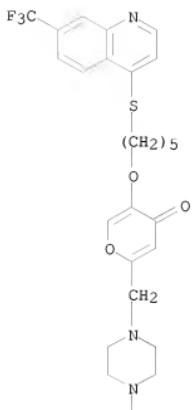
RN 754240-25-0 HCPLUS

CN 4H-Pyran-4-one, 2-[[4-(methylsulfonyl)-1-piperazinyl]methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]thio]pentyl]oxy]- (CA INDEX NAME)



RN 754240-26-1 HCPLUS

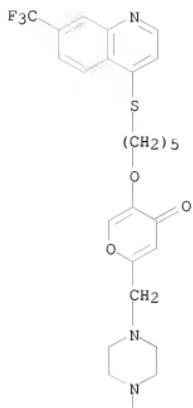
CN 1-Piperazinecarboxamide, N-(1,1-dimethylethyl)-4-[(4-oxo-5-[[5-[(7-(trifluoromethyl)-4-quinolinyl)thio]pentyl]oxy]-4H-pyran-2-yl)methyl]-(CA INDEX NAME)



RN 754240-27-2 HCPLUS

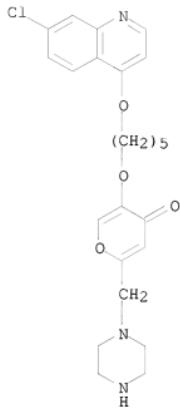
CN 1-Piperazinecarboxamide, N-methyl-4-[[4-oxo-5-[[5-[(7-(trifluoromethyl)-4-

quinolinyl]thiopentyl]oxy]-4H-pyran-2-yl)methyl]- (CA INDEX NAME)



RN 754240-28-3 HCAPLUS

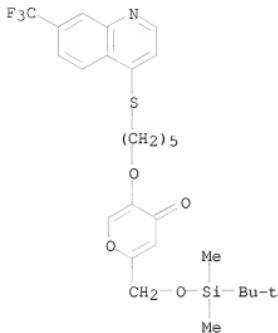
CN 4H-Pyran-4-one, 5-[[5-[(7-chloro-4-quinolinyl)oxy]pentyl]oxy]-2-(1-piperazinylmethyl)-(CA INDEX NAME)



IT 754239-58-2P, 2-[(tert-Butyldimethylsilyloxy)methyl]-5-[(5-[(7-(trifluoromethyl)quinolin-4-yl)sulfonyl]pentyl)oxy]-4H-pyran-4-one  
 754239-61-7P, Methanesulfonic acid [4-oxo-5-[(5-[(7-trifluoromethyl)quinolin-4-ylsulfonyl]pentyl)oxy]-4H-pyran-2-yl]methyl ester 754239-70-8P, 5-[(5-[(6-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-2-(hydroxymethyl)-4H-pyran-4-one 754239-71-9P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-2-(hydroxymethyl)-4H-pyran-4-one 754239-72-0P, 4-[(5-[(6-Hydroxymethyl-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-7-trifluoromethylquinoline-3-carboxylic acid ethyl ester 754239-73-1P, 5-[(5-[(8-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-2-(hydroxymethyl)-4H-pyran-4-one 754239-74-2P, 5-[(5-[(Quinazolin-4-yloxy)pentyl)oxy]-2-(hydroxymethyl)-4H-pyran-4-one 754239-75-3P, 5-[(5-[(6-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-76-4P, 5-[(5-[(7-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-77-5P, 4-[(5-[(6-[(Methanesulfonyloxy)methyl]-4-oxo-4H-pyran-3-yl)oxy]pentyl)oxy]-7-trifluoromethylquinoline-3-carboxylic acid ethyl ester 754239-78-6P, 5-[(5-[(8-(Trifluoromethyl)quinolin-4-yloxy)pentyl)oxy]-4-oxo-4H-pyran-2-yl)methyl methanesulfonate 754239-79-7P, 5-[(5-[(Quinazolin-4-yloxy)pentyl)oxy]-4-oxo-4H-pyran-2-yl)methyl methanesulfonate  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
 (intermediate; preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)

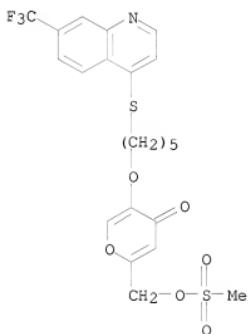
RN 754239-58-2 HCPLUS

CN 4H-Pyran-4-one, 2-[[[(1,1-dimethylethyl)dimethylsilyl]oxy)methyl]-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl)thio]pentyl)oxy]- (CA INDEX NAME)



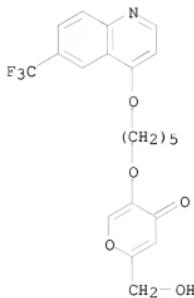
RN 754239-61-7 HCPLUS

CN 4H-Pyran-4-one, 2-[(methylsulfonyl)oxy]methyl-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy]- (CA INDEX NAME)



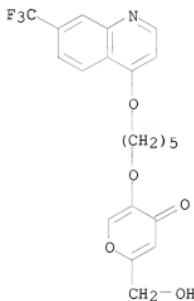
RN 754239-70-8 HCPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[(6-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



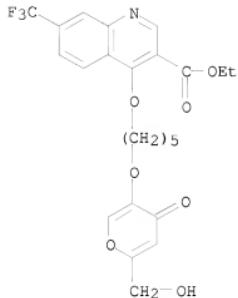
RN 754239-71-9 HCPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]oxy)pentyl]oxy]- (CA INDEX NAME)



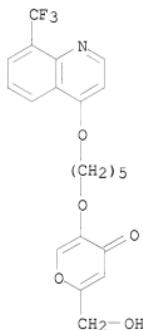
RN 754239-72-0 HCPLUS

CN 3-Quinolinecarboxylic acid, 4-[(5-[(6-(hydroxymethyl)-4-oxo-4H-pyran-3-yl)oxy)pentyl]oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)



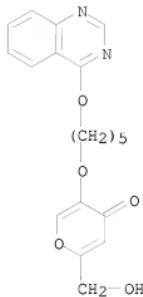
RN 754239-73-1 HCAPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[[5-[[8-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



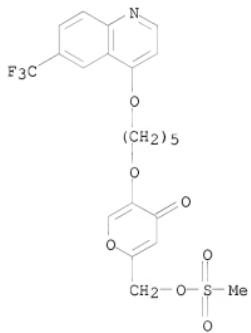
RN 754239-74-2 HCAPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[[5-((4-quinazolinyl)oxy)pentyl]oxy]- (CA INDEX NAME)



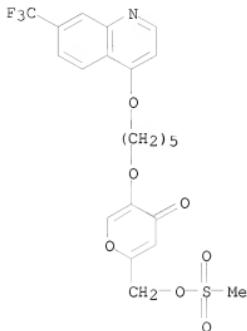
RN 754239-75-3 HCPLUS

CN 4H-Pyran-4-one, 2-[[5-((methylsulfonyl)oxy)methyl]-5-[[5-[[6-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



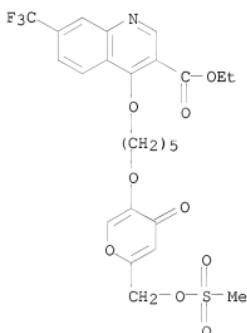
RN 754239-76-4 HCPLUS

CN 4H-Pyran-4-one, 2-[[5-((methylsulfonyl)oxy)methyl]-5-[[5-[[7-(trifluoromethyl)-4-quinolinyl]oxy]pentyl]oxy]- (CA INDEX NAME)



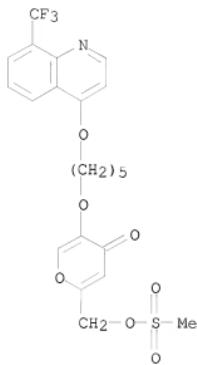
RN 754239-77-5 HCAPLUS

CN 3-Quinolinesulfonyl acid, 4-[[5-[[6-[[((methylsulfonyl)oxy)methyl]-4-oxo-4H-pyran-3-yl]oxy]pentyl]oxy]-7-(trifluoromethyl)-, ethyl ester (CA INDEX NAME)

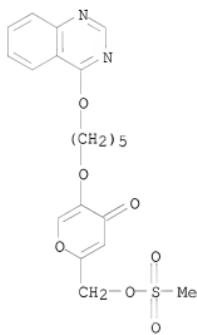


RN 754239-78-6 HCAPLUS

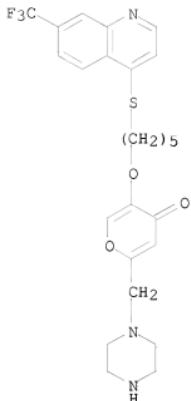
CN 4H-Pyran-4-one, 2-[[((methylsulfonyl)oxy)methyl]-5-[[5-[[8-(trifluoromethyl)-4-quinolinyloxy]pentyl]oxy]- (CA INDEX NAME)



RN 754239-79-7 HCAPLUS  
 CN 4H-Pyran-4-one, 2-[(methylsulfonyl)oxy]methyl-5-[(5-(4-quinazolinyl)oxy)pentyl]oxy- (CA INDEX NAME)



IT 754240-20-5, 5-[(5-[7-(Trifluoromethyl)quinolin-4-ylthio]pentyl)oxy]-2-[(piperazin-1-yl)methyl]-4H-pyran-4-one  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (preparation of novel antiproliferative and antiangiogenic agents, in particular quinoline-derivatized pyranones, for treating cell proliferative diseases, retinopathies and arthritis)  
 RN 754240-20-5 HCAPLUS  
 CN 4H-Pyran-4-one, 2-(1-piperazinylmethyl)-5-[(5-[(7-(trifluoromethyl)-4-quinolinyl]thio)pentyl]oxy)- (CA INDEX NAME)



L12 ANSWER 6 OF 6 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1995:608133 HCPLUS

DOCUMENT NUMBER: 123:83751

ORIGINAL REFERENCE NO.: 123:15005a,15008a

TITLE: Mechanism-Based Development of New Antimalarials:  
Synthesis of Derivatives of Artemisinin Attached to  
Iron Chelators

AUTHOR(S): Kamchonwongpaisan, Sumalee; Paitayatat, Sumpan;  
Thebtaranonth, Yodhathai; Wilairat, Prapin; Yuthavong,  
Yongyuth

CORPORATE SOURCE: Faculty of Science, Mahidol University, Bangkok,  
10400, Thailand

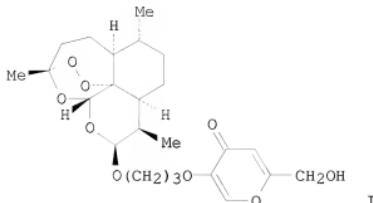
SOURCE: Journal of Medicinal Chemistry (1995), 38(13), 2311-16  
CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

GI



AB Various derivs. of artemisinin covalently linked to iron chelators, e.g. I, were synthesized, and their antimalarial activities were evaluated. Although results show no indication that the presence of an iron chelator in the vicinity of artemisinin potentiates its action, the linked compds. prepared still retain comparable activities to that of artemisinin.

IT 165068-36-0P

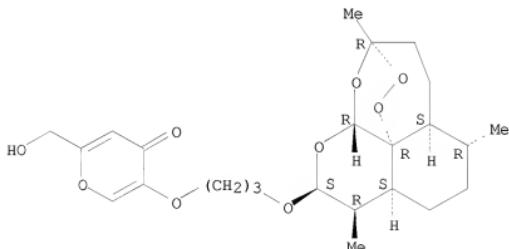
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(mechanism-based development of new antimalarials, synthesis of derivs. of artemisinin attached to iron chelators)

RN 165068-36-0 HCAPLUS

CN 4H-Pyran-4-one, 3-[3-[(decahydro-3,6,9-trimethyl-3,12-epoxy-12H-pyran-4,3-yl)-1,2-benzodioxepin-10-yl]oxylpropoxy]-6-(hydroxymethyl)-, [3R-(3 $\alpha$ ,5 $\alpha$ ,6 $\beta$ ,8 $\beta$ ,9 $\alpha$ ,10 $\alpha$ ,12 $\beta$ ,12aR\*)]- (9CI) (CA INDEX NAME)

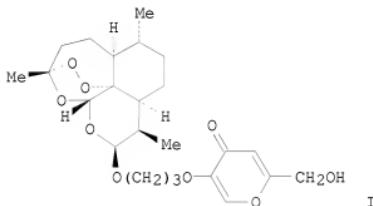
Absolute stereochemistry. Rotation (+).



=> d 114 ibib abs hitstr tot

L14 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2008 ACS on STN  
ACCESSION NUMBER: 1995:608133 HCAPLUS

DOCUMENT NUMBER: 123:83751  
 ORIGINAL REFERENCE NO.: 123:15005a, 15008a  
 TITLE: Mechanism-Based Development of New Antimalarials:  
 Synthesis of Derivatives of Artemisinin Attached to  
 Iron Chelators  
 AUTHOR(S): Kamchonwongpaisan, Sumalee; Paitayatat, Sumparn;  
 Thebtaranonth, Yodhathai; Wilairat, Prapin; Yuthavong,  
 Yongyuth  
 CORPORATE SOURCE: Faculty of Science, Mahidol University, Bangkok,  
 10400, Thailand  
 SOURCE: Journal of Medicinal Chemistry (1995),  
 38(13), 2311-16  
 PUBLISHER: CODEN: JMCMAR; ISSN: 0022-2623  
 DOCUMENT TYPE: American Chemical Society  
 LANGUAGE: Journal  
 English  
 GI



AB Various derivs. of artemisinin covalently linked to iron chelators, e.g. I, were synthesized, and their antimalarial activities were evaluated. Although results show no indication that the presence of an iron chelator in the vicinity of artemisinin potentiates its action, the linked compds. prepared still retain comparable activities to that of artemisinin.

IT 165068-36-0P

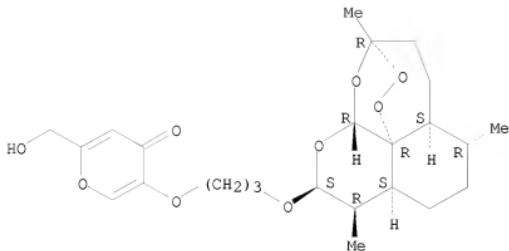
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(mechanism-based development of new antimalarials, synthesis of derivs. of artemisinin attached to iron chelators)

RN 165068-36-0 HCPLUS

CN 4H-Pyran-4-one, 3-[3-[(decahydro-3,6,9-trimethyl-3,12-epoxy-12H-pyran-4,3-dj]-1,2-benzodioxepin-10-yl)oxy]propoxy]-6-(hydroxymethyl)-, [3R-(3a,5aB,6B,8aB,9a,10a,12B,12aR\*)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



=> d l17 ibib abs hitstr tot

L17 ANSWER 1 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:312673 HCPLUS

DOCUMENT NUMBER: 138:321131

TITLE: Preparation of pyranones for treatment of tumors related to GTPases.

INVENTOR(S): Leblanc, Veronique; Leblond, Bertrand; Melle-Milovanovic, Dominique; Lopez Rodriguez, Maria Luz; Viso Beronda, Alma

PATENT ASSIGNEE(S): Exonhit Therapeutics S.A., Fr.

SOURCE: U.S., 24 pp.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

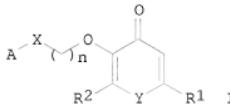
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6552073	B1	20030422	US 2002-85141	20020301 <--
CA 2477003	A1	20030912	CA 2003-2477003	20030228 <--
WO 2003074508	A1	20030912	WO 2003-IB1050	20030228 <--
WO 2003074508	B1	20031127		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 200309924	A1	20030916	AU 2003-209924	20030228 <--
EP 1480966	A1	20041201	EP 2003-743474	20030228
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JP 2005529079	T 20050929	JP 2003-572976	20030228
US 20050054629	A1 20050310	US 2004-502625	20041022 <--
PRIORITY APPLN. INFO.:		US 2002-85141	A 20020301
		WO 2003-IB1050	W 20030228

OTHER SOURCE(S): MARPAT 138:321131  
GI



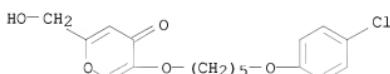
**AB** Title compds. (I; R1 = CH2R3, COR3; R2 = H, alkenyl; R3 = OH, OR4, SR4, NR5R6, pyrrolidinyl, piperidinyl; R4 = alkyl, aryl, aralkyl, alkanoyl, arylcarbonyl; R5, R6 = H, alkyl, aryl, aralkyl; n = 1-10; X = O, S, NR7; Y = O; R7 = H, alkyl, aryl, aralkyl), were prepared. Thus, 5-[7-(3,4-dichloro-2-propylphenoxy)heptyloxy]-2-hydroxymethyl-4H-pyran-4-one (general preparation outlined) showed IC50 = 5-10  $\mu$ M against H460 lung carcinoma and HCT116 colon cancer cells, resp.

**IT**  
 514170-76-4P 514170-77-5P 514170-78-6P  
 514170-79-7P 514170-80-0P 514170-81-1P  
 514170-82-2P 514170-83-3P 514170-84-4P  
 514170-85-5P 514170-86-6P 514170-87-7P  
 514170-88-8P 514170-89-9P 514170-90-2P  
 514170-91-3P 514170-92-4P 514170-93-5P  
 514170-94-6P 514170-95-7P 514170-96-8P  
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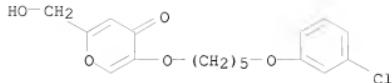
**RL:** PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (claimed compound; preparation of pyranones for treatment of tumors related

**to**  
 GTPases)

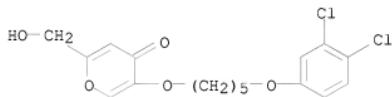
**RN** 514170-76-4 HCPLUS  
**CN** 4H-Pyran-4-one, 5-[5-(4-chlorophenoxy)pentyl]oxy]-2-(hydroxymethyl)- (CA INDEX NAME)



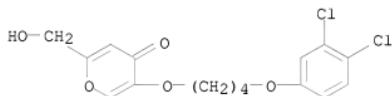
**RN** 514170-77-5 HCPLUS  
**CN** 4H-Pyran-4-one, 5-[5-(3-chlorophenoxy)pentyl]oxy]-2-(hydroxymethyl)- (CA INDEX NAME)



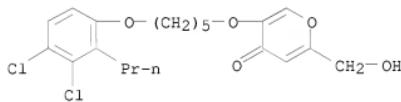
RN 514170-78-6 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[5-(3,4-dichlorophenoxy)pentyl]oxy]-2-(hydroxymethyl)-  
 (CA INDEX NAME)



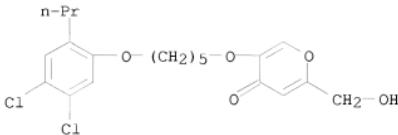
RN 514170-79-7 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[4-(3,4-dichlorophenoxy)butoxy]-2-(hydroxymethyl)-  
 (CA INDEX NAME)



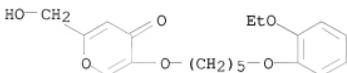
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 (CA INDEX NAME)



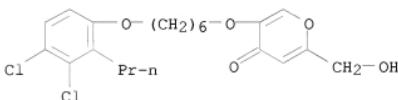
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 (CA INDEX NAME)



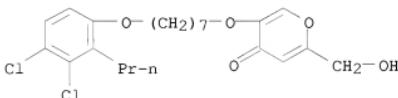
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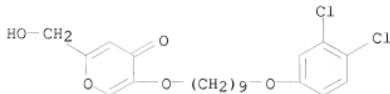
RN 514170-83-3 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[(6-(3,4-dichloro-2-propylphenoxy)hexyl)oxy]-2-(hydroxymethyl)- (CA INDEX NAME)



RN 514170-84-4 HCAPLUS  
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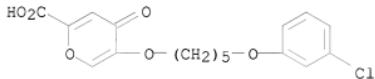
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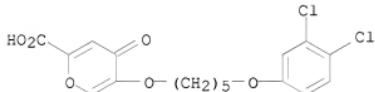
RN 514170-86-6 HCPLUS  
 CN 4H-Pyran-2-carboxylic acid, 5-[(5-(4-chlorophenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)



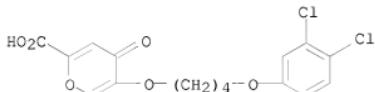
RN 514170-87-7 HCPLUS  
 CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3-chlorophenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)



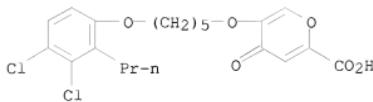
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 CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3,4-dichlorophenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)



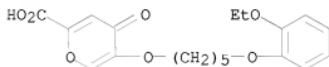
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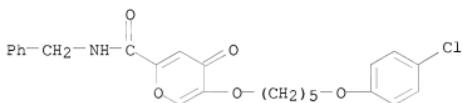
RN 514170-90-2 HCPLUS  
 CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3,4-dichloro-2-propylphenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)



RN 514170-91-3 HCPLUS  
 CN 4H-Pyran-2-carboxylic acid, 5-[(5-(2-ethoxyphenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)

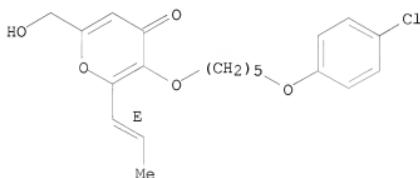


RN 514170-92-4 HCPLUS  
 CN 4H-Pyran-2-carboxamide, 5-[(5-(4-chlorophenoxy)pentyl)oxy]-4-oxo-N-(phenylmethyl)- (CA INDEX NAME)



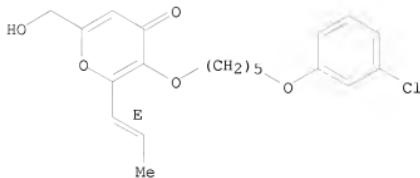
RN 514170-93-5 HCPLUS  
 CN 4H-Pyran-4-one, 3-[(5-(4-chlorophenoxy)pentyl)oxy]-6-(hydroxymethyl)-2-(1E)-1-propen-1-yl- (CA INDEX NAME)

Double bond geometry as shown.



RN 514170-94-6 HCPLUS  
 CN 4H-Pyran-4-one, 3-[(5-(3-chlorophenoxy)pentyl)oxy]-6-(hydroxymethyl)-2-(1E)-1-propen-1-yl- (CA INDEX NAME)

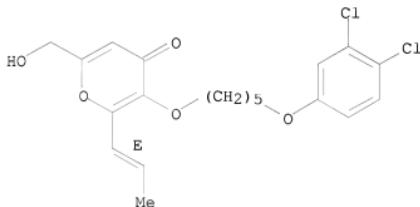
Double bond geometry as shown.



RN 514170-95-7 HCPLUS

CN 4H-Pyran-4-one, 3-[(5-(3,4-dichlorophenoxy)pentyl)oxy]-6-(hydroxymethyl)-2-(1E)-1-propen-1-yl- (CA INDEX NAME)

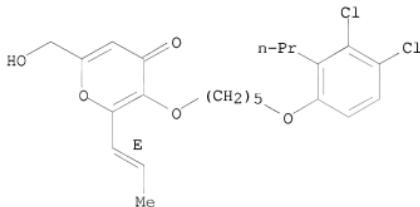
Double bond geometry as shown.



RN 514170-96-8 HCPLUS

CN 4H-Pyran-4-one, 3-[(5-(3,4-dichloro-2-propylphenoxy)pentyl)oxy]-6-(hydroxymethyl)-2-(1E)-1-propen-1-yl- (CA INDEX NAME)

Double bond geometry as shown.

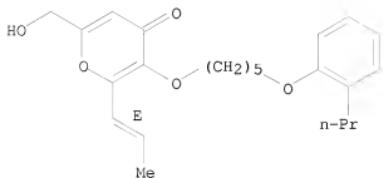


RN 514170-97-9 HCPLUS

CN 4H-Pyran-4-one, 6-(hydroxymethyl)-2-(1E)-1-propen-1-yl-3-[(5-(2-propylphenoxy)pentyl)oxy]- (CA INDEX NAME)

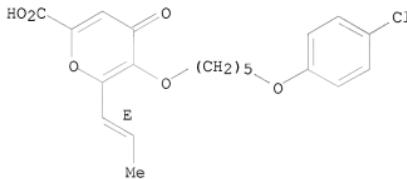
10541328

Double bond geometry as shown.



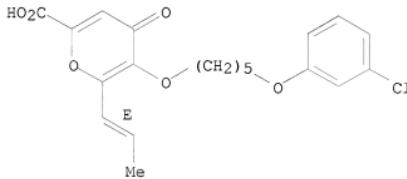
RN 514170-98-0 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 5-[(5-(4-chlorophenoxy)pentyl)oxy]-4-oxo-6-(1E)-1-propen-1-yl- (CA INDEX NAME)

Double bond geometry as shown.



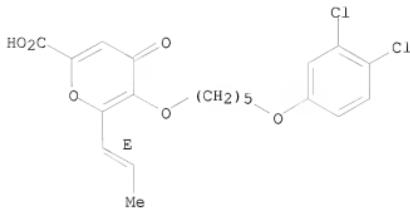
RN 514170-99-1 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3-chlorophenoxy)pentyl)oxy]-4-oxo-6-(1E)-1-propen-1-yl- (CA INDEX NAME)

Double bond geometry as shown.



RN 514171-00-7 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3,4-dichlorophenoxy)pentyl)oxy]-4-oxo-6-(1E)-1-propen-1-yl- (CA INDEX NAME)

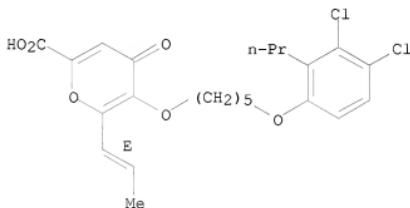
Double bond geometry as shown.



RN 514171-01-8 HCPLUS

CN 4H-Pyran-2-carboxylic acid, 5-[(5-(3,4-dichloro-2-propylphenoxy)pentyl)oxy]-4-oxo-6-(1E)-1-propen-1-yl- (CA INDEX NAME)

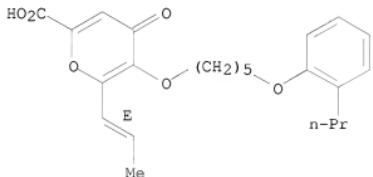
Double bond geometry as shown.



RN 514171-02-9 HCPLUS

CN 4H-Pyran-2-carboxylic acid, 4-oxo-6-(1E)-1-propen-1-yl-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA INDEX NAME)

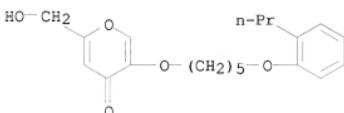
Double bond geometry as shown.



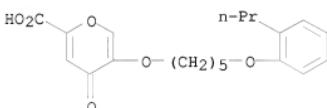
IT 107757-98-2P 107758-01-0P 514171-03-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

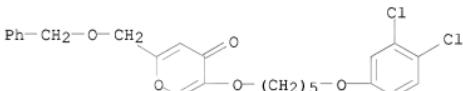
(preparation of pyranones for treatment of tumors related to GTPases)  
 RN 107757-98-2 HCAPLUS  
 CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA  
 INDEX NAME)



RN 107758-01-0 HCAPLUS  
 CN 4H-Pyran-2-carboxylic acid, 4-oxo-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA  
 INDEX NAME)



RN 514171-03-0 HCAPLUS  
 CN 4H-Pyran-4-one, 5-[(5-(3,4-dichlorophenoxy)pentyl)oxy]-2-  
 [(phenylmethoxy)methyl]- (CA INDEX NAME)

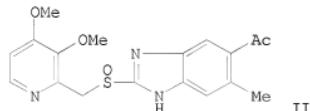
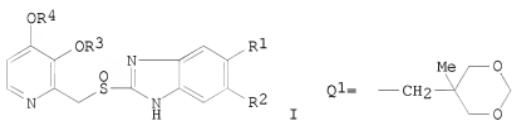


REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS  
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L17 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 1992:151767 HCAPLUS  
 DOCUMENT NUMBER: 116:151767  
 ORIGINAL REFERENCE NO.: 116:25692h,25693a  
 TITLE: Preparation of (pyridinylmethylsulfinyl)benzimidazoles  
 as gastric acid secretion inhibitors  
 Braendstroem, Arne Elof; Lindberg, Per Lennart;  
 Sunden, Gunnar Elisabeth  
 INVENTOR(S): Astra AB, Swed.  
 PATENT ASSIGNEE(S): PCT Int. Appl., 50 pp.  
 SOURCE: CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9119712	A1	19911226	WO 1991-SE416	19910611 <--
W: AT, AU, BB, BG, BR, CA, CH, DE, DK, ES, FI, GB, HU, JP, KP, KR, LK, LU, MC, MG, MW, NL, NO, PL, RO, SD, SE, SU				
RU: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB, GN, GR, IT, LU, ML, MR, NL, SE, SN, TD, TG				
CA 2083606	A1	19911221	CA 1991-2083606	19910611 <--
CA 2083606	C	20010821		
AU 9180617	A	19920107	AU 1991-80617	19910611 <--
AU 649456	B2	19940526		
HU 62862	A2	19930628	HU 1992-4034	19910611 <--
JP 05507714	T	19931104	JP 1991-511436	19910611 <--
JP 3049367	B2	20000605		
EP 593463	A1	19940427	EP 1991-911618	19910611 <--
EP 593463	B1	19990915		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
PL 165898	B1	19950228	PL 1991-297295	19910611 <--
RO 110497	B1	19960130	RO 1992-1543	19910611 <--
AT 184602	T	19991015	AT 1991-911618	19910611 <--
ES 2140391	T3	20000301	ES 1991-911618	19910611 <--
IL 98472	A	19950831	IL 1991-98472	19910612 <--
CN 1058213	A	19920129	CN 1991-105025	19910620 <--
CZ 279434	B6	19950412	CZ 1991-1893	19910620 <--
US 5430042	A	19950704	US 1991-718188	19910620 <--
NO 9204650	A	19921202	NO 1992-4650	19921202 <--
LV 10269	B	19951020	LV 1993-820	19930630 <--
PRIORITY APPLN. INFO.:			SE 1990-2206	A 19900620
			SE 1990-2207	A 19900620
			WO 1991-SE416	A 19910611

OTHER SOURCE(S): MARPAT 116:151767  
GI



AB Title compds. (I; R1, R2 = H, alkyl, COR5; R3, R4 = Me, Et, cyclopropylmethyl, MeOCH2CH2, Q1; R3R4 = (CH2)n; n = 1-3; R5 = alkyl, alkoxy; one of R1, R2 is always COR5) were prepared. Thus,

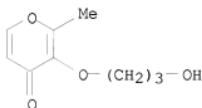
5-acetyl-6-methyl-2-mercaptop-1H-benzimidazole, 3,4-dimethoxy-2-chloromethylpyridine, NaOH, and H<sub>2</sub>O were refluxed in EtOH to give 62% coupling product, which was oxidized with 3-ClC<sub>6</sub>H<sub>4</sub>(O)OOH/NaHCO<sub>3</sub> in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O to give title compound II. II inhibited gastric acid secretion in dogs with ED<sub>50</sub> = 0.74  $\mu$ mol/kg i.v. Tablets were prepared containing II.

IT 139645-13-9P 139645-14-0P 139645-15-1P

RL: SPN (Synthetic preparation); PRP (Preparation)  
(preparation of, as intermediate for ulcer inhibitor)

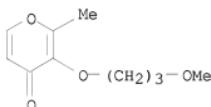
RN 139645-13-9 HCPLUS

CN 4H-Pyran-4-one, 3-(3-hydroxypropoxy)-2-methyl- (CA INDEX NAME)



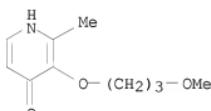
RN 139645-14-0 HCPLUS

CN 4H-Pyran-4-one, 3-(3-methoxypropoxy)-2-methyl- (CA INDEX NAME)



RN 139645-15-1 HCPLUS

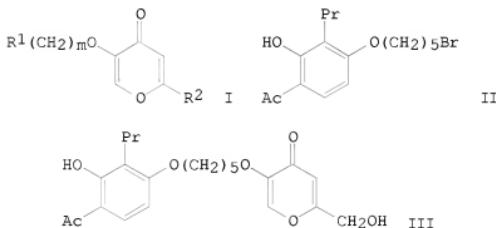
CN 4(1H)-Pyridinone, 3-(3-methoxypropoxy)-2-methyl- (CA INDEX NAME)



L17 ANSWER 3 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 1987:156276 HCPLUS  
 DOCUMENT NUMBER: 106:156276  
 ORIGINAL REFERENCE NO.: 106:25429a,25432a  
 TITLE: Preparation of aralkoxy- and aryloxyalkoxy-substituted  
       kojic acid derivatives as LTD4 inhibitors  
 INVENTOR(S): Masateru, Miyano; Shone, Robert L.  
 PATENT ASSIGNEE(S): G.D. Searle and Co., USA  
 SOURCE: U.S., 11 pp.  
 DOCUMENT TYPE: CODEN: USXXAM  
 Patent

LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 4644071	A	19870217	US 1984-629916	19840711 <--
US 4705871	A	19871110	US 1986-894591	19860808 <--
US 4812584	A	19890314	US 1987-77362	19870724 <--
PRIORITY APPLN. INFO.:			US 1984-629916	A3 19840711
			US 1986-894591	A3 19860808
OTHER SOURCE(S):		CASREACT 106:156276; MARPAT 106:156276		
GI				



AB Title derivs. I [R1 = naphthyl, naphthoxy, 1,2,3,4-tetrahydronaphth-1-or -2-yl, (un)substituted PhO; R2 = CH2OH, CHO, alkoxycarbonyl, CO2H and its alkali-metal or ammonium salts; m = 1-10] are prepared as LTD4 inhibitors (no data). A mixture of 33.2 mmol (bromopentoxy)acetophenone derivative II (preparation given), 26.6 mmol kojic acid, and 60 mmol anhydrous K2CO3

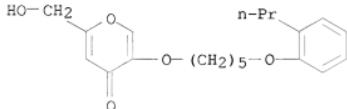
in DMF was stirred at room temperature for 3 days to give 3.0 g of (phenoxypentoxo)pyranone derivative III, a preferred compound

IT 107757-98-2P 107757-99-3P 107758-00-9P  
 107758-01-0P 107758-02-1P 107758-03-2P  
 107758-04-3P 107758-05-4P 107759-81-9P

RL: SPN (Synthetic preparation); PREP (Preparation)  
 (preparation of, as LTD4 inhibitor)

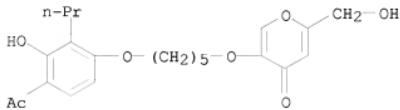
RN 107757-98-2 HCAPLUS

CN 4H-Pyran-4-one, 2-(hydroxymethyl)-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA INDEX NAME)

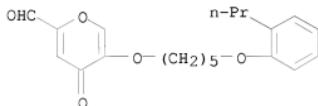


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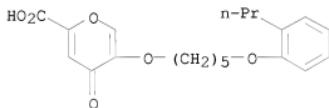
RN 107757-99-3 HCAPLUS  
CN 4H-Pyran-4-one, 5-[(5-(4-acetyl-3-hydroxy-2-propylphenoxy)pentyl)oxy]-2-(hydroxymethyl)- (CA INDEX NAME)



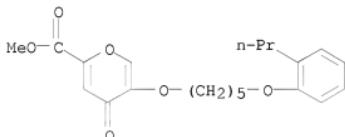
RN 107758-00-9 HCAPLUS  
CN 4H-Pyran-2-carboxaldehyde, 4-oxo-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA INDEX NAME)



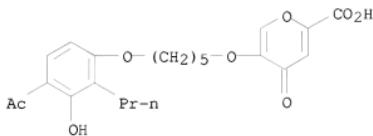
RN 107758-01-0 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 4-oxo-5-[(5-(2-propylphenoxy)pentyl)oxy]- (CA INDEX NAME)



RN 107758-02-1 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 4-oxo-5-[(5-(2-propylphenoxy)pentyl)oxy]-, methyl ester (CA INDEX NAME)

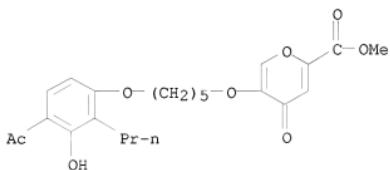


RN 107758-03-2 HCAPLUS  
CN 4H-Pyran-2-carboxylic acid, 5-[(5-(4-acetyl-3-hydroxy-2-propylphenoxy)pentyl)oxy]-4-oxo- (CA INDEX NAME)



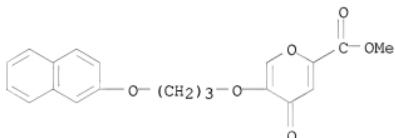
RN 107758-04-3 HCAPLUS

CN 4H-Pyran-2-carboxylic acid, 5-[(5-(4-acetyl-3-hydroxy-2-propylphenoxy)pentyl]oxy]-4-oxo-, methyl ester (CA INDEX NAME)



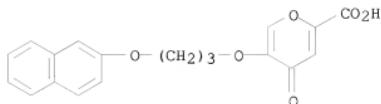
RN 107758-05-4 HCAPLUS

CN 4H-Pyran-2-carboxylic acid, 5-[(3-(2-naphthalenyloxy)propoxy)-4-oxo-, methyl ester (CA INDEX NAME)



RN 107759-81-9 HCAPLUS

CN 4H-Pyran-2-carboxylic acid, 5-[(3-(2-naphthalenyloxy)propoxy)-4-oxo-, sodium salt (9CI) (CA INDEX NAME)



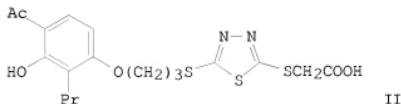
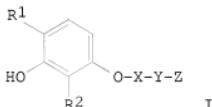
● Na

L17 ANSWER 4 OF 5 HCPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 1986:515074 HCPLUS  
 DOCUMENT NUMBER: 105:115074  
 ORIGINAL REFERENCE NO.: 105:18634h,18635a  
 TITLE: Heterocyclic compounds and their use  
 INVENTOR(S): Kiyoshi, Murase; Toshiyasu, Mase; Hiromu, Hara;  
 Kenichi, Tomioka  
 PATENT ASSIGNEE(S): Yamanouchi Pharmaceutical Co., Ltd., Japan  
 SOURCE: Eur. Pat. Appl., 94 pp.  
 CODEN: EPXWDW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 181779	A1	19860521	EP 1985-308227	19851112 <--
EP 181779	B1	19940316		
R: AT, BE, CH, DE, FR, GB, IT, LI, NL, SE				
ZA 8508493	A	19860730	ZA 1985-8493	19851105 <--
ES 548772	A1	19870901	ES 1985-548772	19851111 <--
SU 1470186	A3	19890330	SU 1985-3979033	19851111 <--
JP 62174057	A	19870730	JP 1985-253562	19851112 <--
JP 63035626	B	19880715		
CA 1269982	A1	19900605	CA 1985-495080	19851112 <--
AT 102930	T	19940415	AT 1985-308227	19851112 <--
ES 556746	A1	19871216	ES 1986-556746	19860625 <--
ES 556747	A1	19871216	ES 1986-556747	19860625 <--
ES 556748	A1	19871216	ES 1986-556748	19860625 <--
ES 556749	A1	19871216	ES 1986-556749	19860625 <--
ES 556750	A1	19871216	ES 1986-556750	19860625 <--
ES 556751	A1	19871216	ES 1986-556751	19860625 <--
ES 556752	A1	19871216	ES 1986-556752	19860625 <--
ES 556753	A1	19871216	ES 1986-556753	19860625 <--
SU 1438610	A3	19881115	SU 1986-4028334	19861021 <--
SU 1491337	A3	19890630	SU 1986-4028351	19861024 <--
SU 1498389	A3	19890730	SU 1986-4028367	19861024 <--
SU 1452481	A3	19890115	SU 1986-4028381	19861027 <--
SU 1493105	A3	19890707	SU 1986-4028350	19861027 <--
SU 1454249	A3	19890123	SU 1986-4028405	19861029 <--
US 4855310	A	19890808	US 1988-173734	19880325 <--
US 4908368	A	19900313	US 1989-362959	19890607 <--

US 5177215	A 19930105	US 1990-486550	19900228 <--
US 5258395	A 19931102	US 1992-960125	19921013 <--
PRIORITY APPLN. INFO.:			
		JP 1984-238991	A 19841112
		JP 1985-219327	A 19851001
		US 1985-796628	A2 19850811
		EP 1985-308227	A 19851112
		US 1988-173734	A3 19880325
		US 1989-362959	A3 19890607
		US 1990-486550	A3 19900228

OTHER SOURCE(S): CASREACT 105:115074; MARPAT 105:115074  
GI



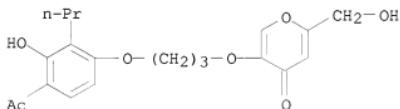
AB Heterocyclic compds. [I; R1 = C1-6 acyl; R2 = C1-6 alkyl; X = C1-6 alkylene or hydroxyalkylene; Y = O, S, carbonylimino, iminocarbonyl; Z = (un)substituted 5- or 6-membered heterocyclic ring containing O, S, N which may be fused with a benzene ring], useful for the treatment and prevention of allergic diseases, are prepared. Thus, thiadiazole II was prepared by reacting 4-(3-bromopropoxy)-2-hydroxy-3-propylacetophenone and [(5-mercapto-1,3,4-thiadiazol-2-yl)thio]acetic acid. II antagonized the actions of SRS-A in guinea pig ileum and trachea and inhibited SRS-A-mediated anaphylactic asthma in conscious guinea pigs. A tablet was formulated containing II 30, lactose 104, starch 57, hydroxypropyl cellulose 4, Ca CM-cellulose 4, and Mg stearate 1 mg.

IT 104073-60-1P

RL: SPN (Synthetic preparation); PREP (Preparation)  
(preparation of, as antiallergic)

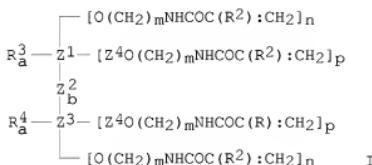
RN 104073-60-1 HCAPLUS

CN 4H-Pyran-4-one, 5-[3-(4-acetyl-3-hydroxy-2-propylphenoxy)propoxy]-2-(hydroxymethyl)- (CA INDEX NAME)



L17 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 1986:461480 HCAPLUS  
 DOCUMENT NUMBER: 105:61480  
 ORIGINAL REFERENCE NO.: 105:10055a,10058a  
 TITLE: Unsaturated cyclic amido-substituted ether compounds  
 INVENTOR(S): Itoh, Hiroshi; Tanaka, Tomio; Nitta, Atsuhiko; Kamio, Hideo  
 PATENT ASSIGNEE(S): Mitsui Toatsu Chemicals, Inc. , Japan  
 SOURCE: Eur. Pat. Appl., 78 pp.  
 CODEN: EPXXDW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

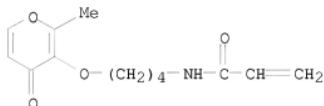
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 155177	A2	19850918	EP 1985-301695	19850312 <--
EP 155177	A3	19870204		
EP 155177	B1	19901031		
R: CH, DE, FR, GB, IT, LI, NL, SE				
JP 60193955	A	19851002	JP 1984-46532	19840313 <--
JP 05073739	B	19931015		
US 4649219	A	19870310	US 1985-708568	19850306 <--
CA 1244012	A1	19881101	CA 1985-475877	19850306 <--
PRIORITY APPLN. INFO.:			JP 1984-46532	A 19840313
OTHER SOURCE(S):	CASREACT 105:61480; MARPAT 105:61480			
GI				



AB Unsatd. cyclic amido-substituted ether compds. I (Z1, Z3 = cyclic group; Z4 = C1-5 alkylene, C2-5 alkenylene, oxyalkylene group, or aminoalkylene group; R3, R4 = halogen, OH, oxo, CN, NO2, SH, S, or a salt thereof, C1-20 alkyl, C2-15 alkenyl, C1-20 haloalkyl, amine group or substituted amine, H, lower alkyl, a carbonyl group, an acid group or salt thereof, or amidopolymethylene group; Z2 = 0, carbonyl, thio, sulfonyl, azo, C1-5 alkylene, C2-5 alkenylene; R2 = H or Me; a = 0-5; m = 4-20; n, p = 0-4 and cannot = 0 at the same time, b = 0 or 1) are useful as crosslinking agents or reactive diluents for hygroscopic polymers. Thus, 1.96 g acrylic amide and 4.0 g 2,2-bis[4-(4-acrylamidobutoxy)phenyl]propane were dissolved in 20 mL DMF and heated at 0-5° for 6 h in the presence of KOH and phenothiazine to give 3.01 g 2,2-bis[4-(4-acrylamidobutoxy)phenyl]propane II. N-Acryloylpyrrolidine containing 0.2% II was mixed with 1% tert-butylperoxy-2-ethylhexanoate and polymerized at 40° for 50 h to give a hygroscopic flexible block polymer.

10541328

IT 102414-01-7P  
RL: PREP (Preparation)  
(preparation of, as crosslinking agent for unsatd. polymers)  
RN 102414-01-7 HCAPLUS  
CN 2-Propenamide, N-[4-[(2-methyl-4-oxo-4H-pyran-3-yl)oxy]butyl]- (CA INDEX  
NAME)



=> log y			
COST IN U.S. DOLLARS		SINCE FILE	TOTAL
		ENTRY	SESSION
FULL ESTIMATED COST		76.16	657.99
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)		SINCE FILE	TOTAL
		ENTRY	SESSION
CA SUBSCRIBER PRICE		-9.60	-13.60

STN INTERNATIONAL LOGOFF AT 10:50:17 ON 04 AUG 2008